LOGINID: sssptau125rxt PASSWORD: TERMINAL (ENTER 1, 2, 3, OR ?):2 Welcome to STN International NEWS Web Page URLs for STN Seminar Schedule - N. America Apr 08 NEWS "Ask CAS" for self-help around the clock NEWS Jun 03 New e-mail delivery for search results now available NEWS Aug 08 PHARMAMarketLetter(PHARMAML) - new on STN NEWS 5 Aug 19 Aquatic Toxicity Information Retrieval (AQUIRE) now available on STN NEWS Aug 26 Sequence searching in REGISTRY enhanced NEWS 7 Sep 03 JAPIO has been reloaded and enhanced NEWS 8 Sep 16 Experimental properties added to the REGISTRY file NEWS 9 Sep 16 CA Section Thesaurus available in CAPLUS and CA NEWS 10 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985 Oct 24 BEILSTEIN adds new search fields NEWS 11 NEWS 12 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN NEWS 13 Nov 18 DKILIT has been renamed APOLLIT NEWS 14 Nov 25 More calculated properties added to REGISTRY NEWS 15 Dec 04 CSA files on STN NEWS 16 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date NEWS 17 Dec 17 TOXCENTER enhanced with additional content NEWS 18 Dec 17 Adis Clinical Trials Insight now available on STN NEWS 19 Jan 29 Simultaneous left and right truncation added to COMPENDEX, ENERGY, INSPEC NEWS 20 Feb 13 CANCERLIT is no longer being updated NEWS 21 Feb 24 METADEX enhancements NEWS 22 Feb 24 PCTGEN now available on STN Feb 24 TEMA now available on STN NEWS 23 NEWS 24 Feb 26 NTIS now allows simultaneous left and right truncation NEWS 25 Feb 26 PCTFULL now contains images NEWS 26 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results NEWS 27 Mar 19 APOLLIT offering free connect time in April 2003 NEWS 28 Mar 20 EVENTLINE will be removed from STN NEWS 29 Mar 24 PATDPAFULL now available on STN NEWS 30 Mar 24 Additional information for trade-named substances without structures available in REGISTRY NEWS 31 Mar 24 Indexing from 1957 to 1966 added to records in CA/CAPLUS Display formats in DGENE enhanced Apr 11 MEDLINE Reload Apr 14

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NEWS EXPRESS April 4 CURRENT WINDOWS VERSION IS V6.01a, CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP), AND CURRENT DISCOVER FILE IS DATED 01 APRIL 2003

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FILE 'HOME' ENTERED AT 15:25:42 ON 16 APR 2003

=> file reg

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 15 APR 2003 HIGHEST RN 503084-53-5 DICTIONARY FILE UPDATES: 15 APR 2003 HIGHEST RN 503084-53-5

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELF PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> s name

L1 4 NAME

=> d 11 4

L1 ANSWER 4 OF 4 REGISTRY COPYRIGHT 2003 ACS

RN 50903-99-6 REGISTRY

CN L-Ornithine, N5-[imino(nitroamino)methyl]-, methyl ester (9CI) (CA INDEX NAME)

OTHER NAMES:

CN L-NAME

CN L-NAME

CN N-Nitro-L-arginine methyl ester

CN N.omega.-Nitro-L-arginine methyl ester

CN N.omega.-Nitro-L-arginine methyl ester

CN NAME

CN NG-Nitro-L-arginine Me ester

CN NG-Nitro-L-arginine methyl ester

FS STEREOSEARCH

DR 162715-84-6, 126265-24-5, 189639-12-1

MF C7 H15 N5 O4

```
LC
     STN Files:
                  ADISINSIGHT, ADISNEWS, AGRICOLA, BEILSTEIN*, BIOBUSINESS,
       BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CEN, CHEMCATS, CIN,
       DIOGENES, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, PROMT, RTECS*,
       TOXCENTER, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
Absolute stereochemistry.
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
            1368 REFERENCES IN FILE CA (1962 TO DATE)
               6 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            1373 REFERENCES IN FILE CAPLUS (1962 TO DATE)
=> s mifepristone
1.2
             5 MIFEPRISTONE
=> d 12 5
L2
     ANSWER 5 OF 5 REGISTRY COPYRIGHT 2003 ACS
     84371-65-3 REGISTRY
RN
     Estra-4,9-dien-3-one, 11-[4-(dimethylamino)phenyl]-17-hydroxy-17-(1-
CN
     propynyl)-, (11.beta., 17.beta.)- (9CI) (CA INDEX NAME)
OTHER NAMES:
     17.beta.-Hydroxy-11.beta.-[4-(dimethylamino)-phenyl]-17.alpha.-(prop-1-
CN
     ynyl)-estra-4,9-dien-3-one
CN
     CDB 2477
CN
     Mifegyne
CN
     Mifeprex
CN
     Mifepristone
CN
     Mifestone
CN
     R 38486
CN
     RU 38486
CN
     RU 486
CN
     RU 486-6
CN
     RU486
FS
     STEREOSEARCH
     122742-25-0, 83203-42-3
DR
MF
     C29 H35 N O2
CI
     COM
LC
                 ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CBNB,
       CEN, CHEMCATS, CIN, CSCHEM, DDFU, DIOGENES, DRUGNL, DRUGPAT, DRUGU,
       DRUGUPDATES, EMBASE, GMELIN*, HSDB*, IPA, MEDLINE, MRCK*, PHAR, PROMT,
       RTECS*, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
```

Absolute stereochemistry.

Other Sources:

CI

COM

## \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

```
1811 REFERENCES IN FILE CA (1962 TO DATE)
59 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
1813 REFERENCES IN FILE CAPLUS (1962 TO DATE)
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```
=> s org31710
L3
             0 ORG31710
=> s org 3171-
           546 ORG
           590 3171
L4
             0 ORG 3171-
                  (ORG(W)3171)
=> s org 31710
           546 ORG
            20 31710
L5
             1 ORG 31710
                  (ORG(W)31710)
=> s 15
           546 ORG
            20 31710
L6
             1 ORG 31710
                  (ORG(W)31710)
=> s 15
           546 ORG
            20 31710
L7
             1 ORG 31710
                  (ORG(W)31710)
=> d 15
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
L5
RN
     118968-41-5 REGISTRY
     Spiro[estra-4,9-diene-17,2'(3'H)-furan]-3-one, 11-[4-
     (dimethylamino)phenyl]-4',5'-dihydro-6-methyl-,
     (6.beta., 11.beta., 17.beta.) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Spiro[17H-cyclopenta[a]phenanthrene-17,2'(3'H)-furan],
     spiro[estra-4,9-diene-17,2'(3'H)-furan]-3-one deriv.
OTHER NAMES:
CN
     Org 31710
FS
     STEREOSEARCH
```

```
MF C30 H39 N O2
CI COM
SR CA
LC STN Files: ADISINSIGHT, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS,
CASREACT, CHEMINFORMRX, DDFU, DRUGUL, DRUGUL, DRUGUPDATES, EMBASE,
MEDLINE, PHAR, TOXCENTER, USPAT2, USPATFULL
```

Absolute stereochemistry.

Absolute stereochemistry.

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

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33 REFERENCES IN FILE CA (1962 TO DATE)
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1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

33 REFERENCES IN FILE CAPLUS (1962 TO DATE)

```
=> s org 33628
           546 ORG
            18 33628
L8
             1 ORG 33628
                  (ORG(W) 33628)
=> d 18
L8
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
RN
     155768-17-5 REGISTRY
     19,24-Dinorchola-4,9,20-trien-3-one, 11-(4-acetylphenyl)-17,23-epoxy-,
     (11.beta., 17.alpha.) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Spiro[17H-cyclopenta[a]phenanthrene-17,2'(3'H)-furan],
     19,24-dinorchola-4,9,20-trien-3-one deriv.
OTHER NAMES:
CN
     Org 33628
FS
     STEREOSEARCH
MF
     C30 H34 O3
SR
     CA
LC
     STN Files:
                  BIOSIS, CA, CAPLUS, DRUGNL, DRUGUPDATES, TOXCENTER, USPATFULL
```

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

13 REFERENCES IN FILE CA (1962 TO DATE)

13 REFERENCES IN FILE CAPLUS (1962 TO DATE)

```
=> s j 867
         69368 J
          1170 867
L9
             1 J 867
                  (J(W)867)
=> d 19
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
L9
RN
     199396-76-4 REGISTRY
     Benzaldehyde, 4-[(11.beta.,17.beta.)-17-methoxy-17-(methoxymethyl)-3-
CN
     oxoestra-4,9-dien-11-yl]-, 1-oxime, [C(E)]- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     Asoprisnil
CN
     J 867
FS
     STEREOSEARCH
DR
     222732-64-1
MF
     C28 H35 N O4
CI
     COM
SR
     CA
LC
     STN Files:
                  ADISINSIGHT, CA, CAPLUS, CASREACT, DRUGUPDATES, PHAR,
```

Absolute stereochemistry. Rotation (+). Double bond geometry as shown.

SYNTHLINE, TOXCENTER, USPATFULL

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

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15 REFERENCES IN FILE CA (1962 TO DATE)
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15 REFERENCES IN FILE CAPLUS (1962 TO DATE)

=> s cdb 2914

57 CDB 534 2914

L10 1 CDB 2914

(CDB(W)2914)

=> d 110

L10 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS

RN 126784-99-4 REGISTRY

CN 19-Norpregna-4,9-diene-3,20-dione, 17-(acetyloxy)-11-[4-(dimethylamino)phenyl]-, (11.beta.)- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN CDB 2914

CN RTI 3021-012

FS STEREOSEARCH

MF C30 H37 N O4

SR CA

LC STN Files: ADISINSIGHT, BEILSTEIN\*, BIOSIS, CA, CANCERLIT, CAPLUS, CASREACT, DRUGNL, DRUGUPDATES, MEDLINE, PROMT, SYNTHLINE, TOXCENTER, USPATFULL

(\*File contains numerically searchable property data)

Absolute stereochemistry.

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

23 REFERENCES IN FILE CA (1962 TO DATE)

23 REFERENCES IN FILE CAPLUS (1962 TO DATE)

=> s zk 137316

248 ZK

2 137316

L11 1 ZK 137316

(ZK(W)137316)

=> d 111

L11 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS

RN 198907-45-8 REGISTRY

CN ZK 137316 (9CI) (CA INDEX NAME)

ENTE A hormonal steroid; an antiprogestin

MF Unspecified

CI MAN SR CA

LC STN Files: BIOSIS, CA, CAPLUS, DRUGNL, DRUGUPDATES, TOXCENTER, USPATFULL

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

12 REFERENCES IN FILE CA (1962 TO DATE)

12 REFERENCES IN FILE CAPLUS (1962 TO DATE)

=> file caplus

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SINCE FILE TOTAL ENTRY SESSION 94.74 94.95

FULL ESTIMATED COST

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FILE COVERS 1907 - 16 Apr 2003 VOL 138 ISS 16 FILE LAST UPDATED: 15 Apr 2003 (20030415/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s nitric oxide synthase

130512 NITRIC

1369614 OXIDE

70286 SYNTHASE

L12 21216 NITRIC OXIDE SYNTHASE

(NITRIC (W) OXIDE (W) SYNTHASE)

=> d his

(FILE 'HOME' ENTERED AT 15:25:42 ON 16 APR 2003)

FILE 'REGISTRY' ENTERED AT 15:25:51 ON 16 APR 2003

L14 S NAME L2 5 S MIFEPRISTONE L3 0 S ORG31710 L40 S ORG 3171-L51 S ORG 31710 L6 1 S L5 L7 1 S L5 Г8 1 S ORG 33628 L9 1 S J 867 L10 1 S CDB 2914 L111 S ZK 137316

FILE 'CAPLUS' ENTERED AT 15:30:05 ON 16 APR 2003

```
L12
          21216 S NITRIC OXIDE SYNTHASE
=> s 11
L13
          1379 L1
=> e fertility
            52
                   FERTILITIES/BI
E2
            1
                   FERTILITTY/BI
E3
         27611 --> FERTILITY/BI
E4
             1
                   FERTILITYARE/BI
E5
             1
                   FERTILITYDISTURBED/BI
Е6
             1
                   FERTILITZER/BI
E7
             3
                   FERTILIY/BI
E8
             8
                   FERTILIZ/BI
E9
             4
                   FERTILIZA/BI
E10
           174
                   FERTILIZABILITY/BI
E11
            82
                   FERTILIZABLE/BI
E12
            1
                   FERTILIZACION/BI
=> s e3
L14
         27611 FERTILITY/BI
=> s 114 and 113
             6 L14 AND L13
=> d 115 1-6
L15 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2003 ACS
     2002:869589 CAPLUS
DN
     Implantation rates after in vitro fertilization, and treatment of
     infertility and early pregnancy loss with a nitric oxide donor or
     substrate alone or in combination with progesterone, and a method for
     contraception with nitric oxide inhibitors in combination with
     antiprogestins or other agents
IN
     Chwalisz, Krzysztof; Garfield, Robert E.
PA
     Germany
     U.S. Pat. Appl. Publ., 15 pp., Division of U.S. Ser. No. 162,446.
SO
     CODEN: USXXCO
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
     -----
                     ____
                           -----
                                          -----
PΙ
    US 2002169205
                      A1
                           20021114
                                          US 2002-43232
                                                           20020114
PRAI US 1998-162446
                      A3
                           19980929
L15
    ANSWER 2 OF 6 CAPLUS COPYRIGHT 2003 ACS
ΑN
    2002:256294 CAPLUS
DN
    136:295093
TI
     Preparation of peptides as melanocortin receptor ligands
    Ebetino, Frank Hallock; Mazur, Adam W.; Hayes, Jeffrey Charles; Wang,
     Feng; Solinsky, Mark Gregory; Colson, Anny-odile; Lin, Qishen
PA
     Procter & Gamble Company, USA
    PCT Int. Appl., 123 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LΑ
    English
FAN.CNT 1
    PATENT NO.
                     KIND DATE
                                          APPLICATION NO.
                                         -----
```

PΙ

WO 2002026774

A2

20020404

WO 2001-US30051 20010926

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W: AE, AG, AL, AM, AT, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EC, EE, EE, ES,
              FI, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG,
             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW,
             MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL,
             TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG,
              KZ, MD, RU, TJ
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI US 2000-235858P
                             20000927
OS
     MARPAT 136:295093
L15 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:128843 CAPLUS
DN
     137:199357
ΤI
     Growth and fertility rates in the offspring of pregnant rats
     treated with L-.omega. nitro-L-arginine methyl ester (L-NAME), a nitric
     oxide inhibitor
AU
     Witlin, Andrea G.; Gangula, Pandu R. R.; Thompson, Mindy L.; Yallampalli,
CS
     Department of Obstetrics and Gynecology, Division of Maternal-Fetal
     Medicine and Division of Reproductive Sciences, The University of Texas
     Medical Branch at Galveston, Galveston, TX, 77555-1062, USA
SO
     American Journal of Obstetrics and Gynecology (2002), 186(1), 89-93
     CODEN: AJOGAH; ISSN: 0002-9378
PB
     Mosby, Inc.
DT
     Journal
LA
     English
RE.CNT 17
               THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 4 OF 6 CAPLUS COPYRIGHT 2003 ACS
AN
     2000:436227 CAPLUS.
DN
     133:291057
ΤI
     Reduction in libido and fertility of male rats by administration
     of the nitric oxide (NO) synthase inhibitor N-nitro-L-arginine methyl
AU
     Ratnasooriya, W. D.; Dharmasiri, M. G.; Wadsworth, R. M.
     Department of Zoology, University of Colombo, Colombo, 3, Sri Lanka
CS
SO
     International Journal of Andrology (2000), 23(3), 187-191
     CODEN: IJANDP; ISSN: 0105-6263
PB
     Blackwell Science Ltd.
DΤ
     Journal
     English
LΑ
RE.CNT 14
               THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 5 OF 6 CAPLUS COPYRIGHT 2003 ACS
L15
     1999:203927 CAPLUS
AN
DN
     131:39874
     Synergistic role of nitric oxide and progesterone during the establishment
ΤI
     of pregnancy in the rat
ΑU
     Chwalisz, Kristof; Winterhager, Elke; Thienel, Thomas; Garfield, Robert E.
CS
     Research Laboratories of Schering AG, Berlin, 13342, Germany
SO
     Human Reproduction (1999), 14(2), 542-552
     CODEN: HUREEE; ISSN: 0268-1161
PB
     Oxford University Press
DT.
     Journal
LΑ
     English
RE.CNT 59
              THERE ARE 59 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
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L15 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2003 ACS
AN
     1994:289794
                 CAPLUS
DN
     120:289794
TΙ
     Effects of nitric oxide-related agents on rat testicular function
     Adams, Michael L.; Meyer, Edward R.; Sewing, Bryan N.; Cicero, Theodore J.
AU
CS
     Sch. Med., Washington Univ., St. Louis, MO, USA
     Journal of Pharmacology and Experimental Therapeutics (1994), 269(1),
SO
     CODEN: JPETAB; ISSN: 0022-3565
DT
     Journal
LΑ
     English
=> d 115 6 all
L15 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2003 ACS
AN
     1994:289794 CAPLUS
DN
     120:289794
ΤI
     Effects of nitric oxide-related agents on rat testicular function
ΑU
     Adams, Michael L.; Meyer, Edward R.; Sewing, Bryan N.; Cicero, Theodore J.
CS
     Sch. Med., Washington Univ., St. Louis, MO, USA
SO
     Journal of Pharmacology and Experimental Therapeutics (1994), 269(1),
     230 - 7
     CODEN: JPETAB; ISSN: 0022-3565
\mathsf{D}\mathbf{T}
     Journal
LΑ
     English
CC
     1-8 (Pharmacology)
     Section cross-reference(s): 2
     The effects of nitric oxide (NO)-related agents on testicular function
AΒ
     were examd. in male rats with measurements of serum LH, serum
     testosterone, testicular interstitial fluid (TIF) testosterone, and TIF
     vols. Serum and TIF testosterone levels and LH secretion were
     significantly decreased by the NO donor, isosorbide dinitrate (ISDN), and
     the NO synthase (NOS) substrate, L-arginine Me ester, a source for the
     endogenous prodn. of NO. The effects of ISDN on TIF vols. were
     inconsistent, but L-arginine Me ester decreased TIF formation in a
     dose-dependent manner. In addn., ISDN dose-dependently suppressed
     testosterone secretion stimulated by human chorionic gonadotropin
     treatment, suggesting that the effects on testosterone secretion were
     independent of changes in secretion of the endogenous gonadotropin LH.
     ISDN, L-arginine Me ester, and the endogenous NOS substrate L-arginine
     completely blocked testosterone secretion stimulated by the NOS inhibitor
     NG-nitro-L-arginine Me ester (NAME), whereas the relatively inactive NOS
     substrate, D-arginine, only partially blocked NAME-stimulated testosterone
     secretion. Hydralazine and nicardipine, two vasodilators that do not
     exhibit prominent NO-related effects, also blocked basal testosterone
     secretion and testosterone secretion stimulated by the vasoconstrictor
     NAME. These results suggest that (1) NO suppresses a major regulatory
     aspect of testicular function, testosterone secretion, (2) the stimulatory
     effects of the NOS inhibitor NAME on testosterone secretion are caused by
     NOS inhibition and a decrease in NO prodn., (3) the vasoactive effects of
     NO and NOS inhibitors, rather than direct steroidogenic effects, may
     mediate these effects on testicular function, and (4) arginine-NOS-NO
     pathways may play an important role in male reproductive endocrine
     function and fertility.
ST
     nitric oxide related agent testicular function
IT
     Testis
        (function, nitric oxide-related agents effect on)
IT
     Vasodilators
        (nitric oxide-related agents as, testicular function response to)
IT
     10102-43-9, Nitric oxide, biological studies
```

```
RL: BIOL (Biological study)
        (agents effect on, testicular function response to)
TT
     125978-95-2, Nitric oxide synthase
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (inhibitor, testicular function response to, nitric oxide role in)
IT
     58-22-0, Testosterone 9002-67-9, LH
     RL: BIOL (Biological study)
        (secretion, nitric oxide-related agents effect on)
IT
     74-79-3, L-Arginine, biological studies 87-33-2, Isosorbide dinitrate
     157-06-2, D-Arginine 2577-94-8, L-Arginine methyl ester
     50903-99-6
     RL: BIOL (Biological study)
        (testicular function response to, nitric oxide role in)
=> d his
   (FILE 'HOME' ENTERED AT 15:25:42 ON 16 APR 2003)
     FILE 'REGISTRY' ENTERED AT 15:25:51 ON 16 APR 2003
              4 S NAME
L1
L2
              5 S MIFEPRISTONE
L3
              0 S ORG31710
L4
              0 S ORG 3171-
L5
              1 S ORG 31710
L6
              1 S L5
L7
              1 S L5
              1 S ORG 33628
L9
              1 S J 867
L10
              1 S CDB 2914
L11
              1 S ZK 137316
     FILE 'CAPLUS' ENTERED AT 15:30:05 ON 16 APR 2003
          21216 S NITRIC OXIDE SYNTHASE
L12
L13
           1379 S L1
                E FERTILITY
L14
          27611 S E3
L15
              6 S L14 AND L13
=> s 114 and 112
L16
           23 L14 AND L12
=> s 116 not 115
L17
           18 L16 NOT L15
=> d 117 1-18
L17 ANSWER 1 OF 18 CAPLUS COPYRIGHT 2003 ACS
AN
    2002:508556 CAPLUS
DN
     137:195831
TI
     Deletion of exon 6 of the neuronal nitric oxide
     synthase gene in mice results in hypogonadism and infertility
ΑU
     Gyurko, Robert; Leupen, Sarah; Huang, Paul L.
CS
    Massachusetts General Hospital and Harvard Medical School, Boston, MA,
    02129, USA
SO
    Endocrinology (2002), 143(7), 2767-2774
    CODEN: ENDOAO; ISSN: 0013-7227
PB
    Endocrine Society
DT
     Journal
LΑ
    English
              THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 35
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
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L17 ANSWER 2 OF 18 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:294227 CAPLUS
DN
     136:315023
ΤI
     Polydithiocarbamate-containing nontargeting macromolecules for therapeutic
     and diagnostic applications
IN
     Lai, Ching-san
PA
     IISA
     U.S. Pat. Appl. Publ., 18 pp., Cont.-in-part of U.S. Ser. No. 899,087,
SO
     abandoned.
     CODEN: USXXCO
DT
     Patent
LА
     English
FAN.CNT 2
     PATENT NO.
                      KIND DATE
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                                           -----
PΙ
     US 2002045573
                      A1
                            20020418
                                           US 1999-409645
                                                            19991001
     CN 1230178
                      Α
                            19990929
                                           CN 1997-197797
     KR 2000035992
                      Α
                            20000626
                                           KR 1999-7001945 19990309
PRAI US 1996-25867P
                      P
                            19960910
     US 1997-899087
                       B2
                            19970723
     US 1996-25867
                       Α
                            19960910
OS
     MARPAT 136:315023
L17 ANSWER 3 OF 18 CAPLUS COPYRIGHT 2003 ACS
     2002:179993 CAPLUS
DN
     137:246002
ΤI
     Nitric oxide synthase and the function of
     sperm in varicocele patients
ΑU
     Zheng, Hang; Zheng, Xinmin; Li, Shiwen
CS
     Center of Urology and Andrology, Second Affiliated Hospital, Wuhan
     university, Wuhan, 430071, Peop. Rep. China
SO
     Zhongguo Nankexue Zazhi (2001), 15(4), 239-241
     CODEN: ZNZHA4; ISSN: 1008-0848
PB
     Shanghai Dier Yike Daxue
DΨ
     Journal
LΑ
     Chinese
L17 ANSWER 4 OF 18 CAPLUS COPYRIGHT 2003 ACS
AN
     2001:577412 CAPLUS
DN
     136:303868
TI
     L-arginine, the substrate of nitric oxide
     synthase, inhibits fertility of male rats
ΑU
     Ratnasooriya, W. D.; Dharmasiri, M. G.
CS
     Department of Zoology, University of Colombo, Colombo, 3, Sri Lanka
     Asian Journal of Andrology (2001), 3(2), 97-103
SO
     CODEN: ASJAF8; ISSN: 1008-682X
PB
     Science Press
DT
     Journal
     English
RE.CNT 21
              THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 5 OF 18 CAPLUS COPYRIGHT 2003 ACS
L17
ΑN
     2001:121007 CAPLUS
DN
     135:75223
     Expression of endothelial nitric oxide
TI
     synthase in the Sertoli cells of men with infertility of various
     causes
ΑU
     Fujisawa, M.; Yamanaka, K.; Tanaka, H.; Tanaka, H.; Okada, H.; Arakawa,
     S.; Kamidono, S.
     Department of Urology, Kobe University School of Medicine, Kobe, Japan
·CS
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- SO BJU International (2001), 87(1), 85-88 CODEN: BJINFO; ISSN: 1464-4096
- PB Blackwell Science Ltd.
- DT Journal
- LA English
- RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 6 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 2001:11841 CAPLUS
- DN 134:113686
- TI Nitric oxide in human sperm
- AU Lewis, Sheena E. M.
- CS Department of Obstetrics and Gynaecology, The Queen's University of Belfast, Belfast, UK
- SO Assisted Reproduction Reviews (1998), 8(2), 58-64 CODEN: AEPEEJ; ISSN: 1051-2446
- PB Decker Periodicals
- DT Journal
- LA English
- RE.CNT 78 THERE ARE 78 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 7 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 2000:774625 CAPLUS
- DN 134:40627
- TI Inducible nitric oxide synthase in the rat testis: Evidence for potential roles in both normal function and inflammation-mediated infertility
- AU O'Bryan, Moira K.; Schlatt, Stefan; Gerdprasert, Orapin; Phillips, David J.; de Kretser, David; Hedger, Mark P.
- CS Monash Inst. Reproduction and Development, Monash Univ., Clayton, 3168, Australia
- SO Biology of Reproduction (2000), 63(5), 1285-1293 CODEN: BIREBV; ISSN: 0006-3363
- PB Society for the Study of Reproduction
- DT Journal
- LA English
- RE.CNT 48 THERE ARE 48 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 8 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 2000:311908 CAPLUS
- DN 133:87259
- TI Activation of platelet-activating factor (PAF) receptor stimulates nitric oxide (NO) release via protein kinase C-.alpha. in HEC-1B human endometrial epithelial cell line
- AU Dearn, Sharon; Rahman, Mabhub; Lewis, Aurelia; Ahmed, Zahra; Eggo, Margaret C.; Ahmed, Asif
- CS Department of Reproductive and Vascular Biology, Division of Reproductive and Child Health, Birmingham Women's Hospital, University of Birmingham, Birmingham, B15 2TG, UK
- SO Molecular Medicine (New York) (2000), 6(1), 37-49 CODEN: MOMEF3; ISSN: 1076-1551
- PB Johns Hopkins University Press
- DT Journal
- LA English
- RE.CNT 44 THERE ARE 44 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 9 OF 18 CAPLUS COPYRIGHT 2003 ACS AN 1999:819234 CAPLUS

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ΤI
     Therapeutic methods employing disulfide derivatives of dithiocarbamates
     and compositions useful therefor
IN
     Lai, Ching-San; Vassilev, Vassil
PA
     Medinox, Inc., USA
SO
     PCT Int. Appl., 102 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                            APPLICATION NO.
                                                              DATE
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                                           WO 1999-US14237 19990622
PI
     WO 9966918
                       A1
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             JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
             TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
             MD, RU, TJ, TM
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     US 6093743
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                                                              19980623
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                             19991229
                       AΑ
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                       Α1
                             20000110
                                            AU 1999-47119
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                                            EP 1999-930617
     EP 1089723
                       Α1
                             20010411
                                                              19990622
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     JP 2002518441
                       Т2
                             20020625
                                            JP 2000-555604
                                                              19990622
     US 6316502
                       В1
                             20011113
                                            US 2000-565666
                                                              20000505
PRAI US 1998-103639
                       A2
                             19980623
     WO 1999-US14237
                       W
                             19990622
     MARPAT 132:59191
OS
RE.CNT 3
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L17
     ANSWER 10 OF 18 CAPLUS COPYRIGHT 2003 ACS
     1999:678176 CAPLUS
AN
     132:48170
DN
ΤI
     Reproductive Function in Female Mice Lacking the Gene for Endothelial
     Nitric Oxide Synthase
     Drazen, Deborah L.; Klein, Sabra L.; Burnett, Arthur L.; Wallach, Edward
ΑU
     E.; Crone, Julie K.; Huang, Paul L.; Nelson, Randy J.
     Behavioral Neuroendocrinology Group, Departments of Psychology and
CS
     Neuroscience, Johns Hopkins University, Baltimore, MD, 21218, USA
     Nitric Oxide (1999), 3(5), 366-374
SO
     CODEN: NIOXF5; ISSN: 1089-8603
PB
     Academic Press
DΤ
     Journal
     English
LΑ
RE.CNT 47
              THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L17
    ANSWER 11 OF 18 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1999:447987 CAPLUS
DN
     131:255383
ΤI
     Reduced tolerance to acute renal ischemia in mice with a targeted
     disruption of the osteopontin gene
ΑU
    Noiri, Eisei; Dickman, Kate; Miller, Frederick; Romanov, Galina; Romanov,
    Victor I.; Shaw, Robert; Chambers, Ann F.; Rittling, Susan R.; Denhardt,
     David T.; Goligorsky, Michael S.
CS
    Department of Medicine, The University of Tokyo, Tokyo, Japan
```

DN

132:59191

- SO Kidney International (1999), 56(1), 74-82 CODEN: KDYIA5; ISSN: 0085-2538
- PB Blackwell Science, Inc.
- DT Journal
- LA English
- RE.CNT 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 12 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 1998:788750 CAPLUS
- DN 130:33045
- TI Method using a nitric oxide scavenger for in vivo reduction of nitric oxide levels, and compositions useful therefor
- IN Lai, Ching-San
- PA MCW Research Foundation, USA
- SO U.S., 20 pp., Cont.-in-part of U.S. Ser. No. 554,196. CODEN: USXXAM
- DT Patent
- LA English
- FAN. CNT 5

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ΡI	US	5847004	Α	19981208	US	1996-767125	19961209		
	US	5756540	Α	19980526	US	1995-459518	19950602		
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	US	6469057	В1	20021022	US	2000-672140	20000927		
	US	2003040511	A1	20030227	US	2002-267528	20021008		
PRAI	US	1995-459518	A2	19950602					
	US	1995-554196	A2	19951106					
	US	1996-767125	A2	19961209					
	US	1997-863059	B2	19970523					
	US	2000-672140	A3	20000927					

- RE.CNT 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD .
  ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 13 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 1998:755828 CAPLUS
- DN 130:151457
- TI Impaired ovulation in mice with targeted deletion of the neuronal isoform of nitric oxide synthase
- AU Klein, Sabra L.; Carnovale, David; Burnett, Arthur L.; Wallach, Edward E.; Zacur, Howard A.; Crone, Julie K.; Dawson, Valina L.; Nelson, Randy J.; Dawson, Ted M.
- CS Behavioral Neuroendocrinology Group, Department of Psychology, Johns Hopkins University, Baltimore, MD, USA
- SO Molecular Medicine (New York) (1998), 4(10), 658-664 CODEN: MOMEF3; ISSN: 1076-1551
- PB Springer-Verlag New York Inc.
- DT Journal
- LA English
- RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L17 ANSWER 14 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 1998:370682 CAPLUS
- DN 129:118187
- TI Chronic blockade of nitric oxide-synthase and endothelin receptors during pregnancy in the rat: effect on pregnancy outcome
- AU Wight, Edward; Kung, Christoph F.; Moreau, Pierre; Takase, Hiroyuki; Luscher, Thomas F.
- CS Department of Obstetrics and Gynecology, University Hospital Zurich,

```
Zurich, CH-8091, Switz.
SO
     Journal of the Society for Gynecologic Investigation (1998), 5(3), 132-139
     CODEN: JSGIED; ISSN: 1071-5576
PB
     Elsevier Science Inc.
DT
     Journal
LΑ
     English
              THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 42
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 15 OF 18 CAPLUS COPYRIGHT 2003 ACS
L17
     1998:180841 CAPLUS
AN
     128:239488
DN
     Polydithiocarbamate-containing macromolecules and the use thereof for
TI
     therapeutic and diagnostic applications
IN
     Lai, Ching-San
PA
     Medinox, Inc., USA; Lai, Ching-San
     PCT Int. Appl., 68 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 2
     PATENT NO. KIND DATE
                                         APPLICATION NO. DATE
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     WO 9811066 A1 19980319
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             PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US,
             US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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     AU 9741725 A1 19980402
                                         AU 1997-41725
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     EP 927159 A1 19990707
                                     EP 1997-939694 19970828
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PRAI US 1996-25867P
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     US 1997-899087
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                      Α
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     WO 1997-US15324
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     MARPAT 128:239488
RE.CNT 1
              THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L17 ANSWER 16 OF 18 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1997:745947 CAPLUS
DN
     128:19047
     Improvement of implantation rates after in vitro fertilization by
     administering a nitric oxide substrate and/or donor
· IN
     Chwalsz, Krzysztof; Garfield, Robert E.
PA
     Schering Aktiengesellschaft, Germany
SO
     PCT Int. Appl., 38 pp.
     CODEN: PIXXD2
DT
     Patent
LΑ
     English
FAN.CNT 1
     PATENT NO.
                    KIND DATE
                                         APPLICATION NO. DATE
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PΙ
     WO 9741866
                       A1
                             19971113
                                            WO 1997-EP2371
                                                             19970507
             AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE,
             ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS,
             LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD,
             SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, UZ, VN, AM, AZ, BY, KG,
             KZ, MD, RU, TJ, TM
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     US 6040340
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                            20000321
                                            US 1996-646518
                                                             19960507
     AU 9728947
                       A1
                             19971126
                                            AU 1997-28947
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     EP 906105
                       A1
                             19990407
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             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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                                            CN 1997-194452
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     BR 9708980
                                            BR 1997-8980
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                                            JP 1997-539553
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     NO 9805204
                       Α
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                                            NO 1998-5204
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PRAI US 1996-646518
                       Α
                            19960507
     WO 1997-EP2371
                       W
                            19970507
L17
     ANSWER 17 OF 18 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1996:324262 CAPLUS
     125:54130
DN
     The immunohistochemical localization of nitric oxide
TT
     synthase (NOS) in the human male reproductive tract and in
     spermatozoa, suggests a possible role for nitric oxide in spermatogenesis
     and sperm maturation and an association with subfertility.
ΑU
     Zini, Armand; O'Bryan, Moira K.; Magid, Margaret; Schlegel, Peter N.
     Department Urology, James Buchanan Brady Foundation, New York, NY, 10021,
CS
SO
     Portland Press Proceedings (1996), 10 (Biology of Nitric Oxide Part 5), 19
     CODEN: POPPEF; ISSN: 0966-4068
PB
     Portland Press
DT
     Journal
     English
LΑ
     ANSWER 18 OF 18 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1995:688648 CAPLUS
     123:80417
DN
TI
     Nitric oxide synthase in the rat fallopian
     tube is regulated during the estrous cycle
ΑU
     Bryant, C. E.; Tomlinson, A.; Mitchell, J. A.; Thiemermann, C.;
     Willoughby, D. A.
     Dep. Exp. Pathol., Med. Coll. St. Bartholomew's Hosp., London, EC1M 6BQ,
CS
     UK
     Journal of Endocrinology (1995), 146(1), 149-57
SO
     CODEN: JOENAK; ISSN: 0022-0795
PB
     Journal of Endocrinology
DT
     Journal
LA
     English
=> d 117 12-18 all
    ANSWER 12 OF 18 CAPLUS COPYRIGHT 2003 ACS
L17
     1998:788750 CAPLUS
ΑN
DN
     130:33045
TI
     Method using a nitric oxide scavenger for in vivo reduction of nitric
     oxide levels, and compositions useful therefor
ΙN
     Lai, Ching-San
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PA MCW Research Foundation, USA
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SO U.S., 20 pp., Cont.-in-part of U.S. Ser. No. 554,196. CODEN: USXXAM

DT Patent

LA English

IC ICM A01N037-18

NCL 514599000

CC 1-12 (Pharmacology)

Section cross-reference(s): 63

## FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO. DATE
PI	US 5847004	Α	19981208	US 1996-767125 19961209
	US 5756540	Α	19980526	US 1995-459518 19950602
	US 5741815	A	19980421	US 1995-554196 19951106
	US 6469057	В1	20021022	US 2000-672140 20000927
	US 2003040511	A1	20030227	US 2002-267528 20021008
PRAI	US 1995-459518	A2	19950602	
	US 1995-554196	A2	19951106	
	US 1996-767125	A2	19961209	
	US 1997-863059	B2	19970523	
	US 2000-672140 ·	A3	20000927	
7 10	Motherda			

Methods are provided for the in vivo redn. of nitric oxide levels in a AB mammalian subject. In contrast to the inhibitory approach described in the prior art (i.e., wherein the function of the enzymes responsible for nitric oxide prodn. is inhibited), the present invention employs a scavenging approach whereby overproduced nitric oxide is bound in vivo to a suitable nitric oxide scavenger. The resulting complex renders the nitric oxide harmless, and is eventually excreted in the urine of the host. An exemplary nitric oxide scavenger contemplated for use in the practice of the present invention is a dithiocarbamate-ferrous iron complex. This complex binds to .NO, forming a stable, water-sol. NO-contg. complex having a characteristic three-line spectrum (indicative of a mononitrosyl-Fe complex) which can readily be detected at ambient temps. by EPR spectroscopy. The invention relates to methods for reducing in vivo levels of .NO as a means of treating subjects afflicted with inflammatory and/or infectious disease. Nitric oxide scavengers are administered to a host in need of such treatment; these scavengers interact with in vivo produced .NO, forming a stable NO-contg. complex. The NO-contg. complex is then filtered through the kidneys, concd. in the urine, and eventually excreted by the subject, thereby reducing in vivo .NO levels.

ST nitric oxide scavenger therapeutic

IT AIDS (disease)

(AIDS dementia complex, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT Mental disorder

(AIDS dementia, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT Intestine, disease

(Crohn's, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT Nervous system

(Huntington's chorea, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT Respiratory distress syndrome

(adult, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT Transplant rejection

(allotransplant, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT Nervous system

(amyotrophic lateral sclerosis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Nervous system Nervous system (central, trauma, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Fatigue, biological . (chronic fatigue syndrome, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Pain (chronic, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) TΤ Nervous system (degeneration, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Mental disorder (depression, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Gastrointestinal motility (disorder, dysmotility, dysmotility, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Sexual behavior (disorder, priapism, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (emulsions; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT **Fertility** (enhancement, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Heart, disease Kidney, disease (failure, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Stomach, disease (gastritis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Kidney, disease (glomerulonephritis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ITTransplant and Transplantation (graft-vs.-host reaction, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IΤ Neoplasm (hematol., nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) TT Dialysis (hemodialysis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Shock (circulatory collapse) (hemorrhagic, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Appetite (hyperphagia, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Intestine, disease (ileitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Intestine (ileum, disease, ileitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level)

ΙT

Liver, disease

Liver, disease (inflammation, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Intestine, disease (inflammatory, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (inhalants; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (injections, i.v.; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (injections, s.c.; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Head Lung, disease Reperfusion (injury, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) Brain, disease ΙT (ischemia, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (liposomes; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (liqs., dispersions; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drugs (lung injury induced by, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Meningitis (lymphocytic choriomeningitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Headache (migraine, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Myeloproliferative disorders (myelofibrosis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Heart, disease (myocarditis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Kidney, disease (nephritis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) TΤ AIDS (disease) Alzheimer's disease Anaphylaxis Anxiety Arthritis Asthma Atherosclerosis Autoimmune disease Burn Cachexia Cardiopulmonary bypass Cirrhosis Cystic fibrosis Dermatitis

Diabetes mellitus

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Drug dependence
     Eczema
     Encephalomyelitis
     Epilepsy
     Heart, disease
     Hepatitis
     Infection
     Inflammation
     Ischemia
     Liver, disease
     Malaria
     Meningitis
     Multiple sclerosis
     Neoplasm
     Obesity
     Parkinson's disease
     Psoriasis
     Schizophrenia
     Transplant rejection
     Ulcer
     Urticaria
        (nitric oxide overprodn. assocd. with; nitric oxide scavenger for in
        vivo redn. of nitric oxide level)
ΙT
     Cytokines
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); BSU (Biological study, unclassified); THU (Therapeutic use);
     BIOL (Biological study); PROC (Process); USES (Uses)
        (nitric oxide overprodn. assocd. with; nitric oxide scavenger for in
        vivo redn. of nitric oxide level)
IT
     Interleukin 1
     Interleukin 12
     Interleukin 2
     Interleukin 6
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (nitric oxide overprodn. assocd. with; nitric oxide scavenger for in
        vivo redn. of nitric oxide level)
IT
     Interferons
     Interleukins
     Tumor necrosis factors
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (nitric oxide overprodn. assocd. with; nitric oxide scavenger for in
        vivo redn. of nitric oxide level)
IT
    Micelles
     Scavengers
     Spin trapping
        (nitric oxide scavenger for in vivo redn. of nitric oxide level)
IT
     Lipopolysaccharides
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (nitric oxide scavenger for in vivo redn. of nitric oxide level)
TΤ
    Antibiotics
     Cardiovascular agents
        (nitric oxide scavenger for in vivo redn. of nitric oxide level, and
        combination use)
IT
    Catecholamines, biological studies
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (nitric oxide scavenger for in vivo redn. of nitric oxide level, and
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combination use) IT Drug delivery systems (oral; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Pancreas, disease (pancreatitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Drug delivery systems (parenterals; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Peritoneum (peritonitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Ovarian cycle (premenstrual syndrome, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Transplant and Transplantation (preservation, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Drug delivery systems (rectal; nitric oxide scavenger for in vivo redn. of nitric oxide level) ΙT Shock (circulatory collapse) (septic, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) TΤ Neoplasm (solid, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Drug delivery systems (solids; nitric oxide scavenger for in vivo redn. of nitric oxide IT' Drug delivery systems (solns.; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Brain, disease (stroke, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Lupus erythematosus (systemic, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Shock (circulatory collapse) (toxic shock syndrome, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Intestine, disease (ulcerative colitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT Eye, disease (uveitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IΤ Blood vessel, disease (vasculitis, nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IΤ Interferons RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (.gamma., nitric oxide overprodn. assocd. with; nitric oxide scavenger for in vivo redn. of nitric oxide level) IT 160525-37-1 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (nitric oxide scavenger for in vivo redn. of nitric oxide level) IT 151268-43-8

RL: BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence); PROC

(nitric oxide scavenger for in vivo redn. of nitric oxide level)

74-79-3, L-Arginine, biological studies IΤ 125978-95-2, Nitric

oxide synthase

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(nitric oxide scavenger for in vivo redn. of nitric oxide level)

IT 10102-43-9, Nitric oxide, biological studies

RL: BPR (Biological process); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process)

(nitric oxide scavenger for in vivo redn. of nitric oxide level)

ΙT 51-41-2, Noradrenaline 51-61-6, Dopamine, biological studies 34368-04-2, Dobutamine

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(nitric oxide scavenger for in vivo redn. of nitric oxide level, and combination use)

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- L17 ANSWER 13 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN1998:755828 CAPLUS
- DN 130:151457
- Impaired ovulation in mice with targeted deletion of the neuronal isoform ΤI of nitric oxide synthase
- ΑU Klein, Sabra L.; Carnovale, David; Burnett, Arthur L.; Wallach, Edward E.; Zacur, Howard A.; Crone, Julie K.; Dawson, Valina L.; Nelson, Randy J.; Dawson, Ted M.
- CS Behavioral Neuroendocrinology Group, Department of Psychology, Johns Hopkins University, Baltimore, MD, USA

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SO
     Molecular Medicine (New York) (1998), 4(10), 658-664
     CODEN: MOMEF3; ISSN: 1076-1551
PB
     Springer-Verlag New York Inc.
DT
     Journal
LА
     English
CC
     13-6 (Mammalian Biochemistry)
     Section cross-reference(s): 14
AB
     Background: Nitric oxide (NO) plays an important role in numerous
     reproductive processes. To date, most studies have assessed the role of
     NO by using nonspecific pharmacol. inhibitors of the precursor to NO,
     nitric oxide synthase (NOS). These pharmacol.
     NOS inhibitors suppress all isoforms of NOS; thus, the precise
     contribution of each isoform to female reproductive physiol. is unknown.
     The purpose of this study was to det. the specific role of neuronal NOS
     (nNOS) in the regulation of ovulation in female mice lacking the gene that
     encodes for nNOS (nNOS-/-). Materials and Methods: Ovulation was assessed
     in wild-type (WT) and nNOS-/- female mice by examg. the no. of ovarian
     rupture sites and no. of oocytes recovered from the oviducts following
     mating or exposure to exogenous gonadotropins (i.e., 5 IU pregnant mares
     serum gonadotropin [PMSG] and 5 IU human chorionic gonadotropin [hCG]).
     Ovulatory efficiency was detd. as the no. of ovulated oocytes per no. of
     ovarian rupture sites. To examine whether ovulatory deficits in nNOS-/-
     mice were due to alterations in central mechanisms, plasma LH concns. were
     assessed in WT and nNOS-/- mice that were challenged with 25 ng of
     gonadotropin-releasing hormone (GnRH). To det. whether ovulatory deficits
     in nNOS-/- mice were due to local ovulation processes, nerves innervating
     the reproductive tract of WT and nNOS-/- females were examd. for the
     presence of nNOS protein. Results: There were substantial
     fertility deficits in nNOS-/- female mice; the nNOS-/- mice had
     fewer oocytes in their oviducts following spontaneous and
     gonadotropin-stimulated ovulation. Pituitary responsiveness to exogenous
     GnRH challenge was intact in nNOS-/- mice. Dense nNOS protein staining
     was obsd. in nerves innervating the reproductive tracts of WT mice.
     Conclusions: The reproductive deficits in nNOS-/- females are most likely
     due to alterations in the transfer of oocytes from the ovaries to the
     oviducts during ovulation. These results suggest that defects in
     neuronally derived NO prodn. may contribute to female infertility.
ST
     nitric oxide synthase ovulation infertility
     Mutation
IT
        (deletion; impaired ovulation in mice with targeted deletion of
        neuronal isoform of nitric oxide synthase
ΙT
     Fertility
       Fertility
        (female, disorder; impaired ovulation in mice with targeted deletion of
        neuronal isoform of nitric oxide synthase
IT
     Ovulation
        (impaired ovulation in mice with targeted deletion of neuronal isoform
        of nitric oxide synthase)
IT
     10102-43-9, Nitric oxide, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (impaired ovulation in mice with targeted deletion of neuronal isoform
        of nitric oxide synthase)
IT
     125978-95-2, Nitric oxide synthase
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (neuronal isoform; impaired ovulation in mice with targeted deletion of
        neuronal isoform of nitric oxide synthase
RE.CNT
       29
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- L17 ANSWER 14 OF 18 CAPLUS COPYRIGHT 2003 ACS
- AN 1998:370682 CAPLUS
- DN 129:118187
- TI Chronic blockade of nitric oxide-synthase and endothelin receptors during pregnancy in the rat: effect on pregnancy
- Wight, Edward; Kung, Christoph F.; Moreau, Pierre; Takase, Hiroyuki; AU Luscher, Thomas F.
- CS Department of Obstetrics and Gynecology, University Hospital Zurich, Zurich, CH-8091, Switz.
- SO Journal of the Society for Gynecologic Investigation (1998), 5(3), 132-139 CODEN: JSGIED; ISSN: 1071-5576
- Elsevier Science Inc. PΒ
- DTJournal
- LΑ English
- CC 2-10 (Mammalian Hormones)
- To investigate the effects of endothelin-1 (ET-1) receptor antagonism AΒ and/or chronic blockade of nitric oxide (NO) prodn. on pregnancy outcome in the rat. Pregnant or nonpregnant Wistar rats were either treated orally for up to 18 days with the NO-synthase inhibitor N.omega.-nitro-L-arginine Me ester (L-NAME), the ETA-/ETB-receptor antagonist bosentan or both, or received no treatment (controls). pressure, body wt., and drug intake were measured at regular intervals. Pregnancy outcome and proteinuria were also detd. Anal. of variance and paired Student t test were used for statistical anal. Chronic L-NAME treatment increased systolic blood pressure by 69 and 64 mmHg in pregnant and virgin rats resp. Bosentan-blunted, L-NAME-induced hypertension at the beginning, but not at the end of the treatment period in all rats examd. N.omega.-nitro-L-arginine Me ester-treatment in pregnancy reduced the no. of living fetuses at term and caused proteinuria. Bosentan tended to reverse the effects of L-NAME on fetus no. and proteinuria, but both effects failed to reach statistical significance. The effects of chronic

NO-synthase-blockade on blood pressure in gravid rats can be reversed only temporarily by ETA-/ETB-antagonism, suggesting an involvement of endothelin-1 in the early phase of the L-NAME-induced, preeclampsia-like syndrome during pregnancy, although at later stages other mechanisms may come into play. pregnancy nitric oxide endothelin receptor **Fertility** (female; nitric oxide-synthase and endothelin receptor chronic blockade effect on pregnancy outcome in rats) Pregnancy (nitric oxide-synthase and endothelin receptor chronic blockade effect on pregnancy outcome in rats) Endothelin receptors RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (nitric oxide-synthase and endothelin receptor chronic blockade effect on pregnancy outcome in rats) 10102-43-9, Nitric oxide, biological studies 123626-67-5, Endothelin-1 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (nitric oxide-synthase and endothelin receptor chronic blockade effect on pregnancy outcome in rats) 116243-73-3, Endothelin 125978-95-2, Nitric oxide synthase RL: BSU (Biological study, unclassified); BIOL (Biological study) (nitric oxide-synthase and endothelin receptor chronic blockade effect on pregnancy outcome in rats) RE.CNT THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) Baylis, C; Clin Exp Hypertens Pregnancy 1992, VB11, P117 CAPLUS (2) Boulanger, C; Circ Res 1992, V70, P1191 CAPLUS (3) Boulanger, C; J Clin Invest 1990, V85, P587 CAPLUS (4) Bunag, R; Hypertension 1982, V4, P898 MEDLINE (5) Clozel, M; J Pharmacol Exp Ther 1994, V270, P228 CAPLUS (6) Diket, A; Am J Obstet Gynecol 1994, V171, P1243 CAPLUS (7) Furchgott, R; Ann Rev Pharmacol Toxicol 1984, V24, P175 CAPLUS (8) Garg, U; J Clin Invest 1989, V83, P1774 CAPLUS (9) Goerre, S; Circulation 1995, V91, P359 MEDLINE (10) Grieff, M; Transplantation 1993, V56, P880 MEDLINE (11) Helmbrecht, G; Am J Obstet Gynecol 1996, V175, P800 CAPLUS (12) Jaffer, F; Kidney Int 1990, V38, P1193 CAPLUS (13) Joannides, R; Circulation 1995, V91, P1314 CAPLUS (14) Kamoi, K; N Engl J Med 1990, V323, P1486 MEDLINE (15) Konishi, M; Hypertension 1983, V5, P881 CAPLUS (16) Kurihara, Y; Nature 1994, V368, P703 CAPLUS (17) Li, J; Hypertension 1994, V24, P183 CAPLUS (18) Luscher, T; Hypertension 1986, V8(Suppl 2), P55 (19) Luscher, T; J Cardiovasc Pharmacol 1994, V24(Suppl 3), PS16 (20) Luscher, T; The endothelium: Modulator of cardiovascular function 1990 (21) Macgillivray, I; Clin Sci 1969, V37, P395 MEDLINE (22) Magness, R; Semin Perinatol 1991, V15, P68 MEDLINE (23) Masaki, T; Semin Perinatol 1991, V15, P27 MEDLINE (24) Molnar, M; Am J Obstet Gynecol 1994, V170, P1458 CAPLUS (25) Nava, E; Circulation 1995, V91, P2310 CAPLUS (26) Palmer, S; Obstet Gynecol 1992, V80, P1000 MEDLINE (27) Peeters, L; Am J Obstet Gynecol 1980, V138, P1177 MEDLINE (28) Rapoport, R; Circ Res 1983, V52, P352 CAPLUS (29) Rees, D; Br J Pharmacol 1990, V101, P746 CAPLUS (30) Richard, V; Circulation 1995, V91, P771 CAPLUS (31) Schiff, E; Am J Obstet Gynecol 1992, V166, P624 MEDLINE

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L17
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AN
     1998:180841 CAPLUS
DN
     128:239488
ΤI
     Polydithiocarbamate-containing macromolecules and the use thereof for
     therapeutic and diagnostic applications
ΙN
     Lai, Ching-San
PA
     Medinox, Inc., USA; Lai, Ching-San
     PCT Int. Appl., 68 pp.
SO
     CODEN: PIXXD2
DΤ
     Patent
LΑ
     English
IC
     ICM C07C333-14
CC
     1-12 (Pharmacology)
     Section cross-reference(s): 8
FAN.CNT 2
     PATENT NO.
                        KTND
                              DATE
                                               APPLICATION NO.
PΙ
     WO 9811066
                        A1
                               19980319
                                               WO 1997-US15324 19970828
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
          RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,
              GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
              GN, ML, MR, NE, SN, TD, TG
     AU 9741725
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     AU 746790
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     EP 927159
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                                               EP 1997-939694
                                                                  19970828
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, FI
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     KR 2000035992
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PRAI US 1996-25867P
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     US 1997-899087
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     US 1996-25867
                         Α
                              19960910
     WO 1997-US15324
                         W
                              19970828
os
     MARPAT 128:239488
AΒ
     A new class of drugs is provided for therapeutic treatment of such
     indications as cerebral stroke and other ischemia/reperfusion injury.
     Dithiocarbamates are linked to the surface of a macromol. (e.g. albumin),
     either by using crosslinking reagents or by non-specific binding, to
     produce polydithiocarbamate-macromol.-contg. compns. Combination
     therapeutic methods have been developed for the in vivo inactivation or
     inhibition of formation (either directly or indirectly) of species which
     induce the expression of inducible nitric oxide
     synthase, as well as reducing nitric oxide levels produced as a
     result of NO synthase expression. Magnetic resonance imaging methods have
     been developed for the measurement of cerebral and cardiac blood flow and
     infarct vol. in ischemic stroke or heart attack situations. Such methods
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employ iron-contg. complexes of a compn. comprising a dithiocarbamate and a macromol. as contrast agents. Prepn. of a reaction product of bovine serum albumin with N-methyl-D-glucamine dithiocarbamate is described. ST dithiocarbamate macromol therapeutic diagnostic; albumin dithiocarbamate therapeutic diagnostic; MRI contrast dithiocarbamate macromol iron complex; cerebral stroke ischemia reperfusion dithiocarbamate macromol; nitric oxide synthase dithiocarbamate macromol ΙT AIDS (disease) AIDS (disease) (AIDS dementia complex; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Mental disorder Mental disorder (AIDS dementia; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Nervous system (Huntington's chorea; polydithiocarbamate-contq. macromols. for therapeutic and diagnostic applications) IT Imaging agents Imaging agents (NMR contrast; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Respiratory distress syndrome (adult; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Skin, disease (aging, disorder, photoaging, and photodamage; polydithiocarbamatecontg. macromols. for therapeutic and diagnostic applications) ΙT Transplant rejection (allotransplant; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Nervous system (amyotrophic lateral sclerosis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) ΙT Nervous system Nervous system (central, trauma; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Fatigue, biological (chronic fatigue syndrome; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Nervous system (degeneration; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) Gastrointestinal motility IT (disorder, dysmotility, dysmotility; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) ΙT **Fertility** (enhancement; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Heart, disease Kidney, disease (failure; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT (gastrointestinal; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) ΙT Kidney, disease (glomerulonephritis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Transplant and Transplantation (graft-vs.-host reaction; polydithiocarbamate-contg. macromols. for

therapeutic and diagnostic applications) IT Antitumor agents Antitumor agents (hematol.; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Dialysis (hemodialysis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Shock (circulatory collapse) (hemorrhagic; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Appetite (hyperphagia; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Intestine, disease (ileitis; polydithiocarbamate-contq. macromols. for therapeutic and diagnostic applications) IT Intestine (ileum, disease, ileitis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) ΙT Intestine, disease (inflammatory; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Lung, disease Reperfusion (injury; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Brain, disease (ischemia, focal; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Brain, disease (ischemia; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Infection (leading to overprodn. of inflammatory cytokines; polydithiocarbamatecontg. macromols. for therapeutic and diagnostic applications) IT (lung injury from; polydithiocarbamate-contq. macromols. for therapeutic and diagnostic applications) IT Meningitis (lymphocytic choriomeningitis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Myeloproliferative disorders (myelofibrosis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) ITHeart, disease (myocarditis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Cytoprotective agents (neuroprotectants; polydithiocarbamate-contq. macromols. for therapeutic and diagnostic applications) ITScavengers (nitric oxide; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Pancreas, disease (pancreatitis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Peritoneum (peritonitis; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) IT Skin, disease

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(photoaging, and photodamage; polydithiocarbamate-contg. macromols. for
        therapeutic and diagnostic applications)
ΙT
     Analgesics
     Anaphylaxis
     Anti-AIDS agents
     Anti-Alzheimer's agents
     Anti-inflammatory agents
     Anti-ischemic agents
     Antiarthritics
     Antiasthmatics
     Anticonvulsants
     Antidepressants
     Antidiabetic agents
     Antimalarials
     Antimigraine agents
     Antiobesity agents
     Antiparkinsonian agents
     Antitumor agents
     Antiulcer agents
     Anxiolytics
     Autoimmune disease
     Burn
     Cachexia
     Cardiovascular agents
     Cirrhosis
     Cystic fibrosis
     Dermatitis
     Drug delivery systems
     Drug dependence
     Eczema
     Encephalomyelitis
     Eye, disease
     Hepatitis
     Liver, disease
     Meningitis
     Multiple sclerosis
     Nervous system agents
     Organ preservation
     Psoriasis
     Schizophrenia
     Shock (circulatory collapse)
     Transplant rejection
     Urticaria
        (polydithiocarbamate-contg. macromols. for therapeutic and diagnostic
        applications)
IT
     Cytokines
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (polydithiocarbamate-contg. macromols. for therapeutic and diagnostic
        applications)
ΙT
     Ovarian cycle
        (premenstrual syndrome; polydithiocarbamate-contg. macromols. for
        therapeutic and diagnostic applications)
IT
     Penis
        (priapism; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
IT
    Hemocyanins
    Nucleic acids
    Ovalbumin
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (reaction products with dithiocarbamates; polydithiocarbamate-contg.
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Polysaccharides, biological studies
     Proteins, specific or class
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (reaction products, with dithiocarbamates; polydithiocarbamate-contg.
        macromols. for therapeutic and diagnostic applications)
IT
     Albumins, reactions
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (reaction; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
     Shock (circulatory collapse)
IT
        (septic; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
     Antitumor agents
ΙT
        (solid tumor; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
IT
     Brain, disease
        (stroke; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
     Lupus erythematosus
        (systemic; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
ΙT
     Shock (circulatory collapse)
        (toxic shock syndrome; polydithiocarbamate-contg. macromols. for
        therapeutic and diagnostic applications)
IT
     Bacteria (Eubacteria)
        (translocation; polydithiocarbamate-contg. macromols. for therapeutic
        and diagnostic applications)
IT
     Shock (circulatory collapse)
        (traumatic; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
IT
     Blood vessel, disease
        (vasculitis; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
IT
     125978-95-2, Nitric oxide synthase
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (inducible; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
IT
     7439-89-6, Iron, biological studies
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (overload; polydithiocarbamate-contg. macromols. for therapeutic and
        diagnostic applications)
ΙT
     594-07-0D, Dithiocarbamic acid, dithiocarbamate-macromol. products
     1398-61-4D, Chitin, dithiocarbamate reaction products 7439-89-6D, Iron,
     complexes with dithiocarbamate-macromol. products, biological studies
     9000-07-1D, Carrageenan, dithiocarbamate reaction products
                                                                 9000-30-0D,
     Guar gum, dithiocarbamate reaction products
                                                  9000-40-2D, Locust bean gum,
     dithiocarbamate reaction products
                                        9000-69-5D, Pectin, dithiocarbamate
     reaction products 9004-34-6D, Cellulose, dithiocarbamate reaction
    products, biological studies
                                   9004-54-0D, Dextran, dithiocarbamate
     reaction products, biological studies
                                            9004-61-9D, Hyaluronic acid,
    dithiocarbamate reaction products
                                       9005-25-8D, Starch, dithiocarbamate
     reaction products, biological studies
                                            9005-32-7D, Alginic acid,
    dithiocarbamate reaction products
                                        9005-49-6D, Heparin, dithiocarbamate
    reaction products, biological studies 9005-79-2D, Glycogen,
    dithiocarbamate reaction products, biological studies
    Inulin, dithiocarbamate reaction products
                                               9012-36-6D, Agarose,
    dithiocarbamate reaction products 9012-76-4D, Chitosan, dithiocarbamate
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macromols. for therapeutic and diagnostic applications)

Albumins, biological studies

IT

9013-95-0D, Levan, dithiocarbamate reaction products reaction products 11138-66-2D, Xanthan gum, dithiocarbamate reaction products Polymannuronic acid, dithiocarbamate reaction products 39464-87-4D, Scleroglucan, dithiocarbamate reaction products 71010-52-1D, Gellan gum, dithiocarbamate reaction products 94161-07-6D, albumin reaction products RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) 94161-07-6 RL: RCT (Reactant); RACT (Reactant or reagent) (reaction; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) 10102-43-9, Nitric oxide, biological studies RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (scavenger; polydithiocarbamate-contg. macromols. for therapeutic and diagnostic applications) RE.CNT THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) Demuth; US 5387748 A 1995 CAPLUS L17 ANSWER 16 OF 18 CAPLUS COPYRIGHT 2003 ACS 1997:745947 CAPLUS 128:19047 Improvement of implantation rates after in vitro fertilization by administering a nitric oxide substrate and/or donor Chwalsz, Krzysztof; Garfield, Robert E. Schering Aktiengesellschaft, Germany PCT Int. Appl., 38 pp. CODEN: PIXXD2 Patent English ICM A61K031-565 A61K031-57; A61K031-22; A61K031-195; A61K031-34; A61K031-44 2-3 (Mammalian Hormones) Section cross-reference(s): 63 FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_ ---------------WO 9741866 A1 19971113 WO 1997-EP2371 19970507 AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, UZ, VN, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG US 6040340 20000321 Α US 1996-646518 19960507 AU 9728947 19971126 Α1 AU 1997-28947 19970507 EP 906105 19990407 Α1 EP 1997-923032 19970507 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI CN 1218402 Α 19990602 CN 1997-194452 19970507 BR 9708980 Α BR 1997-8980 19990803 19970507 JP 2000510462 Т2 20000815 JP 1997-539553 19970507 NO 9805204 Α 19990106 NO 1998-5204 19981106 KR 2000010833 Α 20000225 KR 1998-708974 19981106 PRAI US 1996-646518 Α 19960507

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AB
     A method is provided for the improvement of implantation rates and/or
     pregnancy rates in a female mammal, comprising administering to a female
     mammal in whom pregnancy is desired an effective amt. of: (a) a
     nitric oxide synthase substrate, a nitric
     oxide donor, or both, optionally in combination with, (b) a progestin,
     and, (c) optionally, in further combination with an estrogen. A method is
     also provided for fertility control for a female mammal,
     comprising administering to a female mammal in whom pregnancy is not
     desired and at risk of becoming pregnant an effective amt. of
     nitric oxide synthase inhibitor in combination
     with an antiprogestin. Pharmaceutical compns. are also provided.
ST
     implantation in vitro fertilization nitric oxide; contraceptive
     nitric oxide synthase inhibitor antiprogestin
IT
     Progestogens
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (antiprogestins; fertility control using a nitric
        oxide synthase inhibitor in combination with an
        antiprogestin)
IT
     Fertility
        (female, disorder; improvement of implantation rates after in vitro
        fertilization by administering a nitric oxide substrate and/or donor)
ΙT
        (female, female fertility disorders; improvement of
        implantation rates after in vitro fertilization by administering a
        nitric oxide substrate and/or donor)
TT
     Contraceptives
        (fertility control using a nitric oxide
        synthase inhibitor in combination with an antiprogestin)
ΙT
     Embryo, animal
        (improvement of implantation rates after in vitro fertilization by
        administering a nitric oxide substrate and/or donor)
IT
     Estrogens
     Progestogens
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (improvement of implantation rates after in vitro fertilization by
        administering a nitric oxide substrate and/or donor and optionally a
        progestin and estrogen)
IT
     Fertilization
        (in vitro; improvement of implantation rates after in vitro
        fertilization by administering a nitric oxide substrate and/or donor)
ΙT
     Pregnancy
        (rate; improvement of implantation rates after in vitro fertilization
        by administering a nitric oxide substrate and/or donor)
IT
     Abortion
        (spontaneous; prevention of early pregnancy loss by administering a
        nitric oxide substrate and/or donor)
IT
     79-17-4, Aminoguanidine
                              504-29-0, 2-Aminopyridine
                                                           1121-58-0,
     4-Methylaminopyridine
                           5407-87-4, 4,6-Dimethyl-2-aminopyridine
     17035-90-4
                  36889-13-1
                               52450-18-7, AMT
                                                53774-63-3
                                                              80471-63-2,
     Epostane
              84371-65-3, Mifepristone
                                          118968-41-5, ORG 31710
     126784-99-4, CDB2914
                            155768-17-5, ORG 33628
                                                     198907-45-8, ZK 137316
     199396-76-4, J 867
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (fertility control using a nitric oxide
        synthase inhibitor in combination with an antiprogestin)
    55-63-0, Nitroglycerin 74-79-3, L-Arginine, biological studies
IΤ
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10102-43-9D, Nitric oxide, substrates and 87-33-2, Isosorbide dinitrate donors 14402-89-2, Sodium nitroprusside 16051-77-7, Isosorbide mononitrate 33876-97-0, SIN-1 125978-95-2D, Nitric oxide synthase, substrates and inhibitors RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (improvement of implantation rates after in vitro fertilization by administering a nitric oxide substrate and/or donor) 50-28-2, Estradiol, biological studies 57-83-0, Progesterone, biological 630-56-8, Hydroxyprogesterone caproate 979-32-8, Estradiol valerate 96346-61-1, Onapristone RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (improvement of implantation rates after in vitro fertilization by administering a nitric oxide substrate and/or donor and optionally a progestin and estrogen) L17 ANSWER 17 OF 18 CAPLUS COPYRIGHT 2003 ACS 1996:324262 CAPLUS 125:54130 The immunohistochemical localization of nitric oxide synthase (NOS) in the human male reproductive tract and in spermatozoa, suggests a possible role for nitric oxide in spermatogenesis and sperm maturation and an association with subfertility. Zini, Armand; O'Bryan, Moira K.; Maqid, Margaret; Schlegel, Peter N. Department Urology, James Buchanan Brady Foundation, New York, NY, 10021, USA Portland Press Proceedings (1996), 10 (Biology of Nitric Oxide Part 5), 19 CODEN: POPPEF; ISSN: 0966-4068 Portland Press Journal English 13-1 (Mammalian Biochemistry) Section cross-reference(s): 14 Nitric oxide synthase (NOS) was present in the human male reproductive tract and in spermatozoa. distributed throughout the cytoplasm of Leydig cells and Sertoli cells in normal and subjects with subfertility. The NOS in the epididymis and vas deferens was almost exclusively confined to the epithelium. Thus, the NOS may play an important role in the function of the human male reproductive system. NO synthase male reproductive tract sperm; subfertility NO synthase male reproductive tract Cytoplasm **Epididymis** Vas deferens (localization and role of nitric oxide synthase in human male reproductive tract and in spermatozoa) Sperm (localization and role of nitric oxide synthase in normal and subfertile human male reproductive tract and in spermatozoa) Testis (Leydig cell, localization and role of nitric oxide synthase in human male reproductive tract and in spermatozoa) Testis (Sertoli cell, localization and role of nitric oxide synthase in human male reproductive tract and in spermatozoa) (disorder, subfertility, localization and role of nitric

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oxide synthase in human male reproductive tract and in spermatozoa) ΙT Reproductive tract (male, localization and role of nitric oxide synthase in normal and subfertile human male reproductive tract and in spermatozoa) IT 125978-95-2, Nitric oxide synthase RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (localization and role of nitric oxide synthase in normal and subfertile human male reproductive tract and in spermatozoa) ANSWER 18 OF 18 CAPLUS COPYRIGHT 2003 ACS AN1995:688648 CAPLUS DN 123:80417 TI Nitric oxide synthase in the rat fallopian tube is regulated during the estrous cycle ΑU Bryant, C. E.; Tomlinson, A.; Mitchell, J. A.; Thiemermann, C.; Willoughby, D. A. CS Dep. Exp. Pathol., Med. Coll. St. Bartholomew's Hosp., London, EC1M 6BQ, SO Journal of Endocrinology (1995), 146(1), 149-57 CODEN: JOENAK; ISSN: 0022-0795 PBJournal of Endocrinology DTJournal LΑ English CC 13-6 (Mammalian Biochemistry) Section cross-reference(s): 2, 7 Nitric oxide produced from L-arginine by nitric oxide AB synthase (NOS) acts in a variety of biol. processes via the stimulation of guanylyl cyclase and subsequent elevation of cGMP. Constitutive, calcium-dependent isoforms of NOS are found in endothelial cells (eNOS) and neurons (nNOS), while macrophages express an inducible, calcium-independent isoform (iNOS) in response to the action of certain cytokines or bacterial endotoxin. While the regulation of NOS by exogenous glucocorticoids and steroid hormones as well documented, the effects of endogenous steroid hormones on NOS activity, such as those released during the estrous cycle, is unknown. Here the authors demonstrate, using specific antibodies for eNOS, nNOS and iNOS, the presence of NOS in the epithelium of rat fallopian tubes at proestrus, late proestrus, estrus, metestrus and diestrus. Western blot anal. of rat fallopian tube homogenates revealed a protein band at approx. 125 kDa, which was recognized by antibodies to different isoforms of NOS, but no bands at the expected mol. wts. (eNOS, 140 kDa; nNOS, 160 kDa; iNOS, 135 kDa). NOS activity in fallopian tubes was measured by the conversion of L-[3H] arginine to L-[3H] citrulline. Both calcium-dependent and -independent NOS activities were present. However, in late proestrus when circulating estrogens are low, NOS activity was reduced in comparison to all other stages of the estrous cycle. Thus the authors show that NOS is present in the epithelial lining of the fallopian tube and is recognized at a previously undescribed mol. wt. The changes in NOS activity in these cells during the estrous cycle may modulate tube motility and contribute to successful fertility. ST nitric oxide synthase oviduct; ovarian cycle nitric oxide synthase ΤT Ovarian cycle (nitric oxide synthase isoforms in rat fallopian tube in relation to estrous cycle) ITEstrogens RL: BAC (Biological activity or effector, except adverse); BSU (Biological

```
study, unclassified); BIOL (Biological study)
        (nitric oxide synthase isoforms in rat
        fallopian tube in relation to estrous cycle)
IT
     Oviduct
        (epithelium, nitric oxide synthase
        isoforms in rat fallopian tube in relation to estrous cycle)
     125978-95-2, Nitric oxide synthase
IT
     RL: BOC (Biological occurrence); BPR (Biological process); BSU (Biological
     study, unclassified); BIOL (Biological study); OCCU (Occurrence); PROC
     (Process)
        (nitric oxide synthase isoforms in rat
        fallopian tube in relation to estrous cycle)
IT
     10102-43-9, Nitric oxide, biological studies
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (nitric oxide synthase isoforms in rat
        fallopian tube in relation to estrous cycle)
=> e implantation
E1
             1
                   IMPLANTATIOH/BI
E2
             1
                   IMPLANTATIOM/BI
         78734 --> IMPLANTATION/BI
E3
Ε4
            27
                   IMPLANTATIONAL/BI
E5
             1
                   IMPLANTATIONAND/BI
             1
E6
                   IMPLANTATIONAT/BI
E7
             1
                   IMPLANTATIONG/BI
           · 2
E8
                   IMPLANTATIONIN/BI
E9
             2
                   IMPLANTATIONM/BI
E10
             2
                   IMPLANTATIONN/BI
E11
             1
                   IMPLANTATIONNALLY/BI
E12
             1
                   IMPLANTATIONON/BI
=> s e3
L18
         78734 IMPLANTATION/BI
=> s 118 and 112
L19
            99 L18 AND L12
=> s 119 and 113
L20
             5 L19 AND L13
=> d 120 1-5
L20 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2003 ACS
AN
     2002:869589 CAPLUS
     137:346927
DN
TI
     Implantation rates after in vitro fertilization, and treatment
     of infertility and early pregnancy loss with a nitric oxide donor or
     substrate alone or in combination with progesterone, and a method for
     contraception with nitric oxide inhibitors in combination with
     antiprogestins or other agents
IN
     Chwalisz, Krzysztof; Garfield, Robert E.
PA
     Germany
SO
     U.S. Pat. Appl. Publ., 15 pp., Division of U.S. Ser. No. 162,446.
     CODEN: USXXCO
DT
     Patent
LΑ
    English
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                             DATE
PΙ
    US 2002169205
                      A1
                            20021114
                                          US 2002-43232
```

- L20 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2003 ACS
- AN 1999:203927 CAPLUS
- DN 131:39874
- TI Synergistic role of nitric oxide and progesterone during the establishment of pregnancy in the rat
- AU Chwalisz, Kristof; Winterhager, Elke; Thienel, Thomas; Garfield, Robert E.
- CS Research Laboratories of Schering AG, Berlin, 13342, Germany
- SO Human Reproduction (1999), 14(2), 542-552 CODEN: HUREEE; ISSN: 0268-1161
- PB Oxford University Press
- DT Journal
- LA English
- RE.CNT 59 THERE ARE 59 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L20 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2003 ACS
- AN 1998:309186 CAPLUS
- DN 129:79672
- TI Chronic nitric oxide synthesis inhibition does not prevent pregnancy vasodilation in the rat
- AU Ahokas, Robert A., Ph. D.; Lubarsky, Suzanne L., M. D.; Park, Gun-Chae, M. D.; Friedman, Steven A., M. D.; Sibai, Baha M., M. D.
- CS Department of Obstetrics and Gynecology, University of Tennessee, Memphis, TN, USA
- SO Hypertension in Pregnancy (1998), 17(1), 55-68 CODEN: HYPPEV; ISSN: 1064-1955
- PB Marcel Dekker, Inc.
- DT Journal
- LA English
- RE.CNT 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L20 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2003 ACS
- AN 1997:572956 CAPLUS
- DN 127:242956
- TI Differential effect of L-NAME and S-methyl-isothiourea on leukocyte emigration in carrageenin-soaked sponge implants in rat
- AU Iuvone, Teresa; Van Osselaer, Nancy; D'acquisto, Fulvio; Carnuccio, Rosa; Herman, Arnold G.
- CS Division of Pharmacology, Faculty of Medicine, University of Antwerpen (UIA), Antwerpen-Wilrijk, B-2610, Belg.
- SO British Journal of Pharmacology (1997), 121(8), 1637-1644 CODEN: BJPCBM; ISSN: 0007-1188
- PB Stockton
- DT Journal
- LA English
- L20 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2003 ACS
- AN 1995:259445 CAPLUS
- DN 122:71778
- TI Inhibitors of **nitric oxide** synthase as potential treatments for opioid withdrawal
- AU London, E. D.; Vaupel, D. B.; Kimes, A. S.
- CS Neuroimaging and Drug Action Section, Natl. Inst. Drug Abuse, Baltimore, MD, USA
- SO Regulatory Peptides (1994), 54(1), 165-6 CODEN: REPPDY; ISSN: 0167-0115
- PB Elsevier
- DT Journal
- LA English

IT

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L20 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2003 ACS
ΑN
     1998:309186 CAPLUS
DN
     129:79672
TΤ
     Chronic nitric oxide synthesis inhibition does not prevent pregnancy
     vasodilation in the rat
     Ahokas, Robert A., Ph. D.; Lubarsky, Suzanne L., M. D.; Park, Gun-Chae, M.
     D.; Friedman, Steven A., M. D.; Sibai, Baha M., M. D.
CS
     Department of Obstetrics and Gynecology, University of Tennessee, Memphis,
     TN, USA
SO
     Hypertension in Pregnancy (1998), 17(1), 55-68
     CODEN: HYPPEV; ISSN: 1064-1955
PB
     Marcel Dekker, Inc.
DT
     Journal
LΑ
     English
CC
     13-6 (Mammalian Biochemistry)
     Section cross-reference(s): 1
     The objective is to det. if blockade of endothelium-derived nitric oxide
AΒ
     synthesis from the day after embryo implantation to the day
     before parturition prevents maternal systemic vasodilation in the rat.
     Timed-pregnant and age-matched nonpregnant Wistar-Kyoto rats were
     administered the nonselective nitric oxide
     synthase inhibitor N.omega.-nitro-L-arginine Me ester (15
     mg/rat/day, s.c.) or saline vehicle (untreated) for 14 days using osmotic
     minipumps. On the last day of treatment (day 20 of gestation in the
     pregnant rats), plasma total nitrate/nitrite concn., mean arterial blood
     pressure, and heart rate were measured. Cardiac output and organ blood
     flows were then measured using radioactive-labeled microspheres for the.
     calcn. of total systemic and organ/tissue vascular conductances, resp.
     Chronic blockade of nitric oxide synthesis decreased plasma
     nitrate/nitrite concn. >90% and induced hypertension with decreased
     cardiac output and organ blood flows in both nonpregnant and pregnant
     rats. Cardiac output and total vascular conductance were significantly
     increased in the pregnant compared to nonpregnant, untreated normotensive
     rats and in nitric-oxide-blocked hypertensive rats. Vascular conductance
     of the skin, skeletal muscle/skeleton, gastrointestinal tract, heart, and
     uterus were significantly greater in pregnant than in nonpregnant rats of
    both treatment groups. Conclusions: Maternal systemic and uterine
     vasodilation during pregnancy is complex and is caused by some
    mechanism(s) other than increased basal endothelium-derived nitric oxide
    prodn. or by a compensatory increase in some other vasodilatory system
    during nitric oxide synthesis blockade.
ST
    nitric oxide inhibition pregnancy vasodilation relationship
IT
     Blood pressure
    Circulation
     Pregnancy
    Vasodilation
        (pregnancy vasodilation independent of chronic nitric oxide synthesis
        inhibition)
IT
    50903-99-6, N.omega.-Nitro-L-arginine methyl ester
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); BIOL (Biological study)
        (pregnancy vasodilation independent of chronic nitric oxide synthesis
       inhibition)
```

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (pregnancy vasodilation independent of chronic nitric oxide synthesis inhibition)

125978-95-2, Nitric oxide synthase

```
10102-43-9, Nitric oxide, biological studies
        RL: BSU (Biological study, unclassified); BIOL (Biological study)
              (pregnancy vasodilation independent of chronic nitric oxide synthesis
              inhibition)
RE.CNT
                        THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Ahokas, R; Am J Obstet Gynecol 1990, V162, P841 MEDLINE
(2) Ahokas, R; Am J Obstet Gynecol 1991, V165, P801 CAPLUS
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(4) Baylis, C; Hypertens Pregn 1994, V13, P342
(5) Buhimschi, I; Human Reprod 1995, V10, P2723 CAPLUS
(6) Conrad, K; Am J Physiol 1989, V257, PR847 CAPLUS
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(9) Danielson, L; Circ Res 1996, V79, P1161 CAPLUS
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(26) Ni, Y; Am J Obstet Gynecol 1997, V176, P856 CAPLUS
(27) Puri, C; Biol Reprod 1982, V27, P967 CAPLUS
(28) Rees, D; Proc Natl Acad Sci USA 1989, V86, P3375 CAPLUS
(29) Slangen, B; Am J Physiol 1996, V270, PH1779 CAPLUS
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(31) Ueda, S; Am J Obstet Gynecol 1986, V155, P195 CAPLUS
(32) Umans, J; Am J Physiol 1990, V259, PF293 CAPLUS
(33) Waldman, S; Pharmacol Rev 1987, V39, P163 CAPLUS
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(35) Weiner, C; Am J Obstet Gynecol 1994, V171, P838 CAPLUS
(36) Weiner, C; Proc Natl Acad Sci USA 1994, V91, P5212 CAPLUS
(37) Whittle, B; Br J Pharmacol 1989, V98, P646 CAPLUS
(38) Yallampalli, C; Am J Obstet Gynecol 1993, V169, P1316 CAPLUS
       ANSWER 4 OF 5 CAPLUS COPYRIGHT 2003 ACS
L20
        1997:572956 CAPLUS
AN
DN
        127:242956
TT
        Differential effect of L-NAME and S-methyl-isothiourea on leukocyte
        emigration in carrageenin-soaked sponge implants in rat
        Iuvone, Teresa; Van Osselaer, Nancy; D'acquisto, Fulvio; Carnuccio, Rosa;
ΑU
        Herman, Arnold G.
CS
        Division of Pharmacology, Faculty of Medicine, University of Antwerpen
        (UIA), Antwerpen-Wilrijk, B-2610, Belg.
SO
        British Journal of Pharmacology (1997), 121(8), 1637-1644 .
        CODEN: BJPCBM; ISSN: 0007-1188
PB
        Stockton
DT
        Journal
LΑ
        English
CC
        1-7 (Pharmacology)
```

The role of nitric oxide (NO) in leukocyte (polymorphonuclear cells,

Section cross-reference(s): 14, 15

ΑB

monocytes and lymphocytes) emigration was studied in a model of carrageenin-sponge implants in rats. The s.c. implantation of 1% (w/v) of .lambda.-carrageenin-soaked sponges elicited an inflammatory response that was characterized by a time-related increase in leukocyte infiltration in the sponges and increased levels of nitrite in the exudate. Total leukocyte infiltration and nitrite prodn. were maximal at 24 h and decreased after 48 and 96 h. The mononuclear cell influx was maximal at 48 h (21% of the total leukocytes). Therefore, this time point was used in the successive expts. Polymorphonuclear cell (PMN) and lymphocyte infiltration in the sponges significantly increased when rats were treated with the non-specific NO-synthase (NOS) inhibitor, NG-nitro-L-arginine methylester (L-NAME) (1 mg ml-1 in drinking water ad libitum). Monocyte emigration was not affected by L-NAME treatment. nitrite levels in the exudate of L-NAME-treated rats were significantly reduced. The concomitant ingestion of L-arginine (30 mg ml-1) resulted in a reversion of the L-NAME effect, while D-arginine (30 mg ml-1) had no effect, indicating the involvement of the L-arginine: NO pathway. Administration of L-NAME resulted also in an increased release of tumor necrosis factor-.alpha. (TNF-.alpha.) and prostacyclin (measured as the stable metabolite, 6-keto-PGF1.alpha.). L-NAME had no effect on monocyte chemoattractant protein-1 (MCP-1) release in the exudate. Since L-NAME may have effects on the local blood flow, phenylephrine (0.034 mg ml-2 in drinking water) was used as it has an effect on the local blood flow similar to L-NAME. Phenylephrine had no effect on either leukocyte emigration, or on nitrite, TNF-.alpha., prostacyclin or MCP-1 accumulation in the exudate. In contrast, the more selective iNOS inhibitor S-methyl-isothiourea (SMT) (10 .mu.g ml-1 in drinking water) significantly reduced PMNs and lymphocyte influx in the sponge, having no effect on monocyte influx. Moreover, SMT decreased nitrite prodn. in the exudate to a comparable extent as L-NAME. Administration of SMT significantly reduced MCP-1 release in the exudate, without an effect on TNF-.alpha. or prostacyclin prodn. Moreover SMT did not produce any changes in local blood flow. The results show that a different outcome of the inflammatory process can be obtained depending on the types of NOS inhibitor used. Furthermore, by inhibiting leukocyte infiltration, the more specific iNOS inhibitors may be useful in the treatment of certain inflammatory reactions which are assocd. with an enhanced formation of NO due to the induction of iNOS.

ST leukocyte emigration inflammation NAME methylisothiourea; nitric oxide synthase inhibitor leukocyte emigration

IT Anti-inflammatory agents

Inflammation

Monocyte

IT

Polymorphonuclear leukocyte

(differential effect of L-NAME and methyl-isothiourea on leukocyte emigration in inflammation from carrageenin-soaked sponge implants in rat in relation to role of nitric oxide and inflammatory mediators) Monocyte chemoattractant protein-1

Tumor necrosis factors

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(differential effect of L-NAME and methyl-isothiourea on leukocyte emigration in inflammation from carrageenin-soaked sponge implants in rat in relation to role of nitric oxide and inflammatory mediators)

IT Cell migration

(leukocyte; differential effect of L-NAME and methyl-isothiourea on leukocyte emigration in inflammation from carrageenin-soaked sponge implants in rat in relation to role of nitric oxide and inflammatory mediators)

IT Cell migration

(lymphocyte; differential effect of L-NAME and methyl-isothiourea on leukocyte emigration in inflammation from carrageenin-soaked sponge

```
implants in rat in relation to role of nitric oxide and inflammatory
        mediators).
ΙT
     Leukocyte
     Lymphocyte
        (migration; differential effect of L-NAME and methyl-isothiourea on
        leukocyte emigration in inflammation from carrageenin-soaked sponge
        implants in rat in relation to role of nitric oxide and inflammatory
        mediators)
ΙT
     2986-19-8, S-Methyl-isothiourea 50903-99-6, NG-Nitro-L-arginine
     methylester,
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (differential effect of L-NAME and methyl-isothiourea on leukocyte
        emigration in inflammation from carrageenin-soaked sponge implants in
        rat in relation to role of nitric oxide and inflammatory mediators)
     10102-43-9, Nitric oxide, biological studies 35121-78-9, Prostacyclin
TΤ
                  125978-95-2, Nitric oxide
     58962-34-8
     synthase
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (differential effect of L-NAME and methyl-isothiourea on leukocyte
        emigration in inflammation from carrageenin-soaked sponge implants in
        rat in relation to role of nitric oxide and inflammatory mediators)
L20 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2003 ACS
AN
    1995:259445 CAPLUS
DN . 122:71778
     Inhibitors of nitric oxide synthase as
     potential treatments for opioid withdrawal
ΑU
     London, E. D.; Vaupel, D. B.; Kimes, A. S.
     Neuroimaging and Drug Action Section, Natl. Inst. Drug Abuse, Baltimore,
CS
    MD, USA
SO
     Regulatory Peptides (1994), 54(1), 165-6
     CODEN: REPPDY; ISSN: 0167-0115
PB
     Elsevier
DT
     Journal
LΑ
    English
CC
     1-11 (Pharmacology)
    The effects of inhibitors of nitric oxide
AΒ
     synthase (NOS) on opioid withdrawal signs e.g. wt. loss, diarrhea,
     wet-dog shakes, etc., were studied in rats that were made
    morphine-dependent by s.c. implantation of morphine pellets.
     The results indicated that the selective brain NOS inhibitor
     7-nitroindazole is more effective in inhibiting the withdrawal signs than
     the 2 nonselective NOS inhibitors, L-NNA and L-NAME.
ST
    nitric oxide synthase inhibitor opioid
    withdrawal
IT
    Brain
    Drug dependence
        (inhibitors of brain nitric oxide synthase
        as potential treatments for opioid withdrawal)
ΙT
    Opioids
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (inhibitors of brain nitric oxide synthase
        as potential treatments for opioid withdrawal)
IT
     57-27-2, Morphine, biological studies
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (inhibitors of brain nitric oxide synthase
        as potential treatments for opioid withdrawal)
ΙT
    2149-70-4, L-NG-Nitroarginine 2942-42-9, 7-Nitroindazole
    50903-99-6, L-NAME
```

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study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (inhibitors of brain nitric oxide synthase
        as potential treatments for opioid withdrawal)
     125978-95-2, Nitric oxide synthase
ΙT
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (inhibitors; inhibitors of brain nitric oxide
        synthase as potential treatments for opioid withdrawal)
=> e contraception
E1
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             2
                   CONTRACEPTING/BI
E3
          2635 --> CONTRACEPTION/BI
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L21
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               CONTRACEPTIV/BI OR CONTRACEPTIVA/BI OR CONTRACEPTIVE/BI OR CONTR
               ACEPTIVELY/BI OR CONTRACEPTIVES/BI)
=> s 121 and 113
L22
             3 L21 AND L13
=> d 122 1-3
L22 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2003 ACS
     2002:869589 CAPLUS
ΑN
DN'
     137:346927
     Implantation rates after in vitro fertilization, and treatment of
TI
     infertility and early pregnancy loss with a nitric oxide donor or
     substrate alone or in combination with progesterone, and a method for
     contraception with nitric oxide inhibitors in combination with
     antiprogestins or other agents
IN
     Chwalisz, Krzysztof; Garfield, Robert E.
PA
     Germany
SO
     U.S. Pat. Appl. Publ., 15 pp., Division of U.S. Ser. No. 162,446.
     CODEN: USXXCO
DТ
     Patent
T.A
     English
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                             DATE
PΙ
    US 2002169205
                      A1
                            20021114
                                           US 2002-43232
                                                             20020114
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RL: BAC (Biological activity or effector, except adverse); BSU (Biological

APPLICATION NO. DATE

CC

FAN.CNT 1

PATENT NO.

2-3 (Mammalian Hormones)

KIND DATE

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PΙ
     WO 9515753
                            19950615
                                           WO 1994-US14133 19941208
                       Α1
         W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB,
             GE, HU, JP, KE, KG, KP, KR, KZ, LK, LT, LU, LV, MD, MG, MN, MW,
             NL, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT, UA, UZ, VN
         RW: KE, MW, SD, SZ, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU,
             MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN,
             TD, TG
     US 5470847
                            19951128
                                           US 1993-165309
                                                            19931210
     AU 9513041
                       A1
                            19950627
                                           AU 1995-13041
                                                            19941208
     US 5643944
                       Α
                            19970701
                                           US 1995-477189
                                                            19950607
     US 5721278
                       Α
                            19980224
                                           US 1995-477187
                                                            19950607
PRAI US 1993-165309
                            19931210
     WO 1994-US14133
                            19941208
AB
     Inhibition of ovulation in a female may be achieved by administering a
     nitric oxide synthase inhibitor, alone or in combination with one or more
     of a progestin, an estrogen, and an LH-RH antagonist, thereby preventing
     conception. The stimulation of ovulation in a female may be achieved by
     administering a nitric oxide source, optionally in further combination
     with one or more of clomiphene, a gonadotropin, and an LH-RH agonist.
     Thus, 27 days old immature rats were injected with 4 IU of pregnant mare's
     serum gonadotropin on day on. Two days later rats were injected with 40
     mg of NG-nitro-L-arginine Me ester at 12 AM and 3 PM and animals were
     sacrificed one day later and examd. for the ovulatory response by counting
     the no. of Graafian follicles 3 and corpora lutea 5 in the ovaries. The
     no. of Graffian follicles and corpora lutea was 9.7 and 0.7 resp. as
     compared to 1.0 and 10.0 for the controls.
ST
     ovulation control nitric oxide synthase inhibition; conception prevention
     nitric oxide synthase inhibition
IT
     Contraceptives
     Insemination, artificial
     Ovarian cycle
     Ovulation
     Pituitary gland
        (ovulation control by regulating nitric oxide levels)
IT
     Estrogens
     Gonadotropins
     Progestogens
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (ovulation control by regulating nitric oxide levels)
IT
     Fertilization
        (extracorporeal, ovulation control by regulating nitric oxide levels)
IT
     Gonadotropins
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (inhibitors, ovulation control by regulating nitric oxide levels)
                     103733-02-4
IT
     9034-40-6, GnRH
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (antagonists; ovulation control by regulating nitric oxide levels)
IT
     50-28-2, 17.beta.-Estradiol, biological studies
                                                       50-50-0, Estradiol
               55-63-0, Nitroglycerin 57-63-6, Ethinyl estradiol
    benzoate
                                                                      57-83-0,
     Progesterone, biological studies
                                       68-23-5, Norethinodrel
                                                                 74-79-3,
     L-Arginine, biological studies 87-33-2, Isosorbide dinitrate
     19-Nortestosterone 520-85-4, Medroxyprogesterone 911-45-5, Clomiphene
                6533-00-2, Norgestrel
     2149-70-4
                                        9002-67-9, LH 9034-40-6D, Lh-rh,
    analogs
              14402-89-2, Sodium nitroprusside 16051-77-7, Isosorbide
    mononitrate
                  17035-90-4
                               17230-88-5, Danazol
                                                     20933-81-7
                                                                   34973-08-5,
    Gonadorelin acetate
                         35189-28-7, Norgestimate 50903-99-6
    54024-22-5, Desogestrel
                                            60282-87-3, Gestodene
                               57444-72-1
                                                                    74381-53-6,
    Leuprolide acetate
                        76932-60-0, Nafarelin acetate
                                                          125978-95-2, Nitric
    oxide synthase 137361-05-8
```

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RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
(ovulation control by regulating nitric oxide levels)
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     FILE 'REGISTRY' ENTERED AT 15:25:51 ON 16 APR 2003
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L2
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L3
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L5
              1 S ORG 31710
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              1 S L5
L7
              1 S L5
rs
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              1 S J 867
L10
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L11
              1 S ZK 137316
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L12
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L13
           1379 S L1
                E FERTILITY
L14
          27611 S E3
              6 S L14 AND L13
L15
L16
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L17
             18 S L16 NOT L15
                E IMPLANTATION
          78734 S E3
L18
L19
             99 S L18 AND L12
L20
              5 S L19 AND L13
                E CONTRACEPTION
L21
          14591 S E3-E10
L22
              3 S L21 AND L13
=> s 12
L23
          1816 L2
=> s 123 and 121
           188 L23 AND L21
L24
=> d 124 170-188
L24
     ANSWER 170 OF 188 CAPLUS COPYRIGHT 2003 ACS
AN
     1988:738 CAPLUS
DN
     108:738
TI
     Effects of RU38486, an antiprogesterone agent, on endometrium, fertilized
     ovum and serum sex steroid levels in rats
ΑU
     Ohtani, Kaori; Sakamoto, Hideki; Takahashi, Toru; Satoh, Nobuyoshi; Den,
     Konbai; Takagi, Shigeo
     Sch. Med., Nihon Univ., Tokyo, Japan
CS
SO
     Nippon Sanka Fujinka Gakkai Zasshi (1987), 39(10), 1709-14
     CODEN: NISFAY; ISSN: 0300-9165
DT
     Journal
LΑ
     Japanese
L24
    ANSWER 171 OF 188 CAPLUS COPYRIGHT 2003 ACS
```

```
AN
     1987:629242 CAPLUS
DN
     107:229242
ΤI
     Contragestion by the progesterone antagonist RU 486: a novel approach to
     human fertility control
ΑU
     Baulieu, Etienne Emile
CS
     Lab. Horm., Fac. Med., Bicetre, 94275, Fr.
SO
     Contraception (1987), 36(Suppl.), 1-5
     CODEN: CCPTAY; ISSN: 0010-7824
DT
     Journal; General Review
LΑ
     English
    ANSWER 172 OF 188 CAPLUS COPYRIGHT 2003 ACS
AN
     1987:547556 CAPLUS
DN
     107:147556
     Interception. III: Postcoital luteal contragestion by an antiprogestin
ΤI
     (mifepristone, RU 486) in 62 women
ΑU
     Van Santen, M. R.; Haspels, A. A.
CS
     Dep. Obstet. Gynecol., Utrecht Univ. Hosp., Utrecht, Neth.
SO
     Contraception (1987), 35(5), 423-31
     CODEN: CCPTAY; ISSN: 0010-7824
DΨ
     Journal
     English
LΑ
L24 ANSWER 173 OF 188 CAPLUS COPYRIGHT 2003 ACS
AN
     1987:527350 CAPLUS
DN
     107:127350
     Interception. IV: Failure of mifepristone (RU 486) as a monthly
     contragestive, "Lunarette"
     Van Santen, M. R.; Haspels, A. A.
ΑU
     Dep. Obstet. Gynecol., Utrecht Univ. Hosp., Utrecht, Neth. Contraception (1987), 35(5), 433-8
CS
SO
     CODEN: CCPTAY; ISSN: 0010-7824
DT
     Journal
LA
     English
L24 ANSWER 174 OF 188 CAPLUS COPYRIGHT 2003 ACS
     1987:509393 CAPLUS
AN
DN
     107:109393
ΤI
     Contragestion by antiprogestin RU 486: a novel approach to human
     fertility control
ΑU
     Baulieu, E. E.; Ulmann, A.; Philibert, D.
CS
     Lab-Hormones, Univ. Paris-Sud, Bicetre, 94275, Fr.
     Serono Symposia Publications from Raven Press (1987), 36(Fertil. Regul.
SO
     Today Tomorrow), 55-73
     CODEN: SPRPDU; ISSN: 0733-897X
DT
     Journal; General Review
LA
     English
L24 ANSWER 175 OF 188 CAPLUS COPYRIGHT 2003 ACS
     1987:423577 CAPLUS
AN
     107:23577
DN
ΤI
    Preparation of estradienolone derivatives useful as antiglucocorticoids
     and antiprogestomimetics, and their pharmaceutical formulation
IN
     Torelli, Vesperto; Teutsch, Jean G.; Philibert, Daniel
PA
     Roussel-UCLAF , Fr.
     U.S., 41 pp. Cont.-in-part of U.S. 4,519,946.
SO
     CODEN: USXXAM
DT
     Patent
LΑ
     English
FAN.CNT 6
     PATENT NO.
                      KIND DATE
                                            APPLICATION NO.
```

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PΙ
     US 4634695
                             19870106
                                             US 1985-693682
                        Α
                                                               19850122
     FR 2497807
                        A1
                             19820716
                                             FR 1981-272
                                                               19810109
     FR 2497807
                        B1
                             19830729
    ·US 4386085
                        Α
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L24
    ANSWER 176 OF 188 CAPLUS COPYRIGHT 2003 ACS
AN
     1987:113695 CAPLUS
DN
     106:113695
     The progesterone antagonist RU 486. A potential new contraceptive
ΤI
     Nieman, Lynnette K.; Choate, Teresa M.; Chrousos, George P.; Healy, David
AU
     L.; Morin, Martin; Renquist, David; Merriam, George R.; Spitz, Irving M.;
     Bardin, C. Wayne; et al.
CS
     Dev. Endocrinol. Branch, Natl. Inst. Child Health Hum. Dev., Bethesda, MD,
SO
     New England Journal of Medicine (1987), 316(4), 187-91
     CODEN: NEJMAG; ISSN: 0028-4793
DT
     Journal
LΑ
     English
L24
     ANSWER 177 OF 188 CAPLUS COPYRIGHT 2003 ACS
AN
     1987:78799 CAPLUS
     106:78799
DN
     Fertility control in women: results with RU 486 by the end of 1985
ΤI
ΑU
     Baulieu, Etienne
CS
     Lab. Horm., INSERM U33, Bicetre, 94275, Fr.
SO
     Journal of Steroid Biochemistry (1986), 25(5B), 847-51
     CODEN: JSTBBK; ISSN: 0022-4731
DT
     Journal; General Review
LΑ
     English
    ANSWER 178 OF 188 CAPLUS COPYRIGHT 2003 ACS
     1986:565206 CAPLUS
AN
     105:165206
DN
     Studies on the antireproductive mechanisms of action of RU 486
TI
     Rojas, Francisco J.; O'Conner, James L.; Asch, Ricardo H.
ÁU
CS
     Health Sci. Cent., Univ. Texas, San Antonio, TX, USA
    Antiprogestin Steroid RU 486 Hum. Fertil. Control, [Proc. Conf. Antiprogestational Compd. RU 486] (1985), Meeting Date 1984, 141-54.
SO
     Editor(s): Baulieu, Etienne-Emile; Segal, Sheldon Jerome. Publisher:
     Plenum, New York, N. Y.
     CODEN: 55DHAY
DΤ
     Conference
LΑ
     English
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L24 ANSWER 179 OF 188 CAPLUS COPYRIGHT 2003 ACS AN 1986:565028 CAPLUS

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DN 105:165028
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- TI RU 486: an antiprogestin steroid with contragestive activity in women
- AU Baulieu, Etienne Emile
- CS Lab. Horm., Univ. Paris Sud, Bicetre, 94270, Fr.
- SO Antiprogestin Steroid RU 486 Hum. Fertil. Control, [Proc. Conf. Antiprogestational Compd. RU 486] (1985), Meeting Date 1984, 1-25. Editor(s): Baulieu, Etienne-Emile; Segal, Sheldon Jerome. Publisher: Plenum, New York, N. Y. CODEN: 55DHAY
- DT Conference; General Review
- LA English
- L24 ANSWER 180 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN . 1986:435637 CAPLUS
- DN 105:35637
- TI Antiprogesterone activity of RU 486 and its contragestive and other applications
- AU Baulieu, E. E.; Ulmann, A.
- CS Fac. Med., INSERM 33, Bicetre, 94275, Fr.
- SO Human Reproduction (1986), 1(2), 107-10 CODEN: HUREEE; ISSN: 0268-1161
- DT Journal; General Review
- LA English
- L24 ANSWER 181 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1986:219236 CAPLUS
- DN 104:219236
- TI Steroid antihormones: the antiprogesterone activity of RU 486 and its contraceptive and other uses.
- AU Baulieu, Etienne Emile; Ulmann, Andre
- CS Univ. Paris-Sud, Paris, Fr.
- SO Bulletin de l'Academie Nationale de Medecine (Paris, France) (1985), 169(8), 1191-9
  CODEN: BANMAC; ISSN: 0001-4079
- DT Journal; General Review
- LA French
- L24 ANSWER 182 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1986:200466 CAPLUS
- DN 104:200466
- TI Modulation of aromatase activity in human endometrial stromal cells by steroids, tamoxifen and RU 486
- AU Tseng, Linda; Mazella, James; Sun, Boling
- CS Sch. Med., State Univ. New York, Stony Brook, NY, 11794, USA
- SO Endocrinology (1986), 118(4), 1312-18 CODEN: ENDOAO; ISSN: 0013-7227
- DT Journal
- LA English
- L24 ANSWER 183 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1986:142404 CAPLUS
- DN 104:142404
- TI The antiprogesterone steroid RU-486 does not impair gonadotropinstimulated luteal adenylyl cyclase activity or gonadotropin release by pituitary cells
- AU Rojas, Francisco J.; O'Conner, James L.; Asch, Ricardo H.
- CS Health Sci. Cent., Univ. Texas, San Antonio, TX, 78284, USA
- SO Journal of Steroid Biochemistry (1985), 23(6A), 1053-8 CODEN: JSTBBK; ISSN: 0022-4731
- DT Journal
- LA English

- L24 ANSWER 184 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1985:589900 CAPLUS
- DN 103:189900
- TI Contragestion by antiprogestin: a new approach to human fertility control
- AU Baulieu, Etienne Emile
- CS Fac. Med. Bicetre, Univ. Paris Sud, Bicetre, 94270, Fr.
- SO Ciba Foundation Symposium (1985), 115 (Abortion), 192-210 CODEN: CIBSB4; ISSN: 0300-5208
- DT Journal; General Review
- LA English
- L24 ANSWER 185 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1985:516457 CAPLUS
- DN 103:116457
- TI Pregnancy prevention by intravaginal delivery of a progesterone antagonist: RU486 tampon for menstrual induction and absorption
- AU Hodgen, Gary D.
- CS Jones Inst. Reprod. Med., East. Virginia Med. Sch., Norfolk, VA, 23507, USA
- SO Fertility and Sterility (1985), 44, 263-7 CODEN: FESTAS; ISSN: 0015-0282
- DT Journal
- LA English
- L24 ANSWER 186 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1985:481878 CAPLUS
- DN 103:81878
- TI The antiprogesterone activity of RU 486, a contragestive agent in the
- AU Sakiz, E.; Euvrard, C.; Baulieu, E. E.
- CS Roussel-Uclaf, Paris, 75007, Fr.
- SO International Congress Series (1984), 655(Endocrinology), 239-42 CODEN: EXMDA4; ISSN: 0531-5131
- DT Journal
- LA English
- L24 ANSWER 187 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1985:215412 CAPLUS
- DN 102:215412
- TI The effects of RU486 on the luteal phase of the rhesus monkey
- AU Asch, Ricardo H.; Rojas, Francisco J.
- CS Health Sci. Cent., Univ. Texas, San Antonio, TX, 78284, USA
- SO Journal of Steroid Biochemistry (1985), 22(2), 227-30 CODEN: JSTBBK; ISSN: 0022-4731
- DT Journal
- LA English
- L24 ANSWER 188 OF 188 CAPLUS COPYRIGHT 2003 ACS
- AN 1984:530975 CAPLUS
- DN 101:130975
- TI Steroid derivatives
- IN Teutsch, Jean G.; Costerousse, Germain; Philibert, Daniel; Deraedt, Roger
- PA Roussel-UCLAF, Fr.
- SO U.S., 33 pp. Cont.-in-part of U.S. 4,386,085. CODEN: USXXAM
- DT Patent
- LA English
- FAN.CNT 6

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	US 4447424 FR 2497807	A A1	19840508 19820716	US 1982-386967 FR 1981-272	19820610 19810109

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FR 2497807
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PRAI FR 1981-272
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L24 ANSWER 188 OF 188 CAPLUS COPYRIGHT 2003 ACS
    1984:530975 CAPLUS
AN
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TI
    Steroid derivatives
IN
    Teutsch, Jean G.; Costerousse, Germain; Philibert, Daniel; Deraedt, Roger
PA
    Roussel-UCLAF , Fr.
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SO U.S., 33 pp. Cont.-in-part of U.S. 4,386,085.
CODEN: USXXAM

DT Patent

LA English

IC A01N045-00; A61K031-56 NCL 424238000

CC 32-5 (Steroids)

Section cross-reference(s): 1, 2

FAN.CNT 6

LAN.	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 4447424	A	19840508	US 1982-386967	19820610
	FR 2497807	A1	19820716	FR 1981-272	19810109
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	US 4519946	Α	19850528	US 1984-614440	19840525
	US 4634695	Α	19870106	US 1985-693682	19850122
	US 4978657	А	19901218 .	US 1985-810316	19851217
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	US 1985-693682		19850122	•	
	US 1985-760703		19850730		
	US 1985-810316		19851217		
GI					•

$$R^4$$
 $R^5$ 
 $R^6$ 
 $R^7$ 
 $R^7$ 
 $R^7$ 
 $R^8$ 
 $R^8$ 

AB Antiglucocorticoid and contraceptive norsteroids I [RR1 = 0, ketal, HON:, CH2:; R = HO, alkoxy, acyloxy, R1 = H; R2R3 = O, bond; R4 = N-, P- or Si-contg. radical, i.e. pyridyl, dimethylaminoalkyl, 4-(Me2NCH2CH2O)C6H4, pyrrolidinophenyl, etc.; R5 = C1-C8 alkyl; R6, R7 = H, HO, alkoxy, acyloxy, HOCH2CO, HO2CCO, alkylcarbamoyl, etc.; R8, R9 =

III

```
cleavage of epoxyestrene derivs. by Grignard reagents. Thus, treatment of
     epoxypropynylestrene II with 4-(Me2N)C6H4MqBr in THF contq. CuBr-Me2S
     complex and subsequent acid hydrolysis gave (aminophenyl)propynylestradien
     e III. At 10 mg/kg/day for 3 days in female rats III inhibited
     implantation 100g, whereas at 500 .mu.g/animal in the rabbit III was
     devoid of progestomimetic activity.
ST
     aminophenylestradienone prepn contraceptive; estradienone
     aminophenyl prepn contraceptive; epoxyestrenol ring cleavage
     Grignard reagent; antiglucocorticoide estradienone
IT
        (by nitrogen-contg. radical substituted estradienones)
ΙT
     Androgens
     Progestogens
     RL: USES (Uses)
        (inhibitors, nitrogen-contg. radical substituted estradienones)
IT
     Contraceptives
        (nitrogen-contg. radical substituted estradienones)
ΙT
     19-Norsteroids
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (prepn. of, of nitrogen-contg. radical substituted estradienones)
     106-95-6, reactions 109-54-6 586-77-6 626-61-9
ΙT
     2052-06-4
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        (Grignard ring cleavage reaction of, with epoxyestrenol deriv.)
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     78-80-8
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        (addn. reaction of, with (aminophenyl)estrenone deriv.)
IT
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (addn. reaction of, with estradienone deriv.)
     5571-36-8
ΙT
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TΨ
     100-61-8, reactions
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        (alkylation of, by isoamyl bromide)
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     91-66-7
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        (bromination of)
     79-01-6, reactions
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        (dechlorination and addn. reaction with (aminophenyl)estrenone deriv.)
ΙT
     91934-73-5
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (epimerization of)
IT
     33403-21-3
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        (epoxide ring cleavage of, with aminophenylmagnesium bromide deriv.)
IT
     90944-65-3P
                   91935-10-3P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
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        (prepn. and Grignard ring cleavage reaction of, with epoxyestrenol
        deriv.)
IT
     91934-77-9P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (prepn. and addn. reaction of, with phenyllithium)
IT
                 91934-81-5P 91934-84-8P 91934-85-9P
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HO, H, alkyl aralkyl; n = 1, 2; optional 16-unsatd.] were prepd. by ring

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91934-86-0P
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        of)
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        (prepn. and bromination of)
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        (substitution reaction of, with bromothiophenol)
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